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Amendments to the Specification:

At page 13, lines 16-30, please replace paragraph with the following:

The Expect value is used as a convenient way to create a significance threshold for reporting results. The default value used for blasting is typically set to 0.0001. In BLAST 2.0, the Expect value is also used instead of the P value (probability) to report the significance of matches. For example, an E value of one assigned to a hit can be interpreted as meaning that in a database of the current size one might expect to see one match with a similar score simply by chance. An E value of zero means that one would not expect to see any matches with a similar score simply by chance. See, e.g., http://www.nebi.nlm.nih.gov/Education/BLASTinfo/. Occasionally, a string of X's or N's will result from a BLAST search. This is a result of automatic filtering of the query for low-complexity sequence that is performed to prevent artifactual hits. The filter substitutes any low-complexity sequence that it finds with the letter "N" in nucleotide sequence (e.g., "NNNNNNNNNNNNNNNN") or the letter "X" in protein sequences (e.g., "XXXXXXXXXXX"). Low-complexity regions can result in high scores that reflect compositional bias rather than significant position-by-position alignment. (Wootton and Federhen, Methods Enzymol 266:554-571, 1996).

At page 23, please replace Table 1J with the following:

Table 1J. BLAST results for NOV1a					
Gene Index/ Identifier	Protein/ Organism	Length (aa)	Identity (%)	Positives (%)	Expect
gi 13569928 ref NP_ 112217.1 NP 112217	a disintegrin-like and metalloprotease with thrombospondin type 1 motif, 12 [Homo sapiens]	1593	269/616 (43%)	371/616 (59%)	e-135
gi 11493589 gb AAG3 5563-1 AF163762_1 (AF163762) AAG35563	zinc metalloendopeptida se [Homo sapiens]	1077	253/624 (40%)	359/624 (56%)	e-128

gi 10645199 ref NP	a disintegrin and	997	247/571	344/571	e-127
	_	997	(43%)	· ·	e-12/
055087.1 	metalloprotease		(436)	(59%)	
NP_055087	with				
	thrombospondin			ļ]
	motifs-7				
	preproprotein; a	ł			
	disintegrin-like		1		
	and			1	1
	metalloprotease				
	(reprolysin type)				
	with				
	thrombospondin				Į
	type 1 motif, 7				}
	[Homo sapiens]				
gi 15309931 ref XP	a disintegrin-like	854	245/604	348/604	e-124
054419.1	and	031	(40%)	(57%)	0 124
XP 054419	metalloprotease		(400)	(3/6/	
AL OSTATS	domain with				
	thrombospondin				
	_				
	type I repeats 10				!
112550001 5122	[Homo sapiens]				
gi 7656869 ref NP 0	a disintegrin and	860	244/630	345/630	e-115
55088.1 NP 055088	metalloprotease		(38%)	(54%)	
	with				
1	thrombospondin		•	:	
1	motifs-6				İ
	preproprotein; a				
	disintegrin-like				
	and				
	metalloprotease			÷	
	(reprolysin type)				
	with				
	thrombospondin				
	type 1 motif, 6				
	[Homo sapiens]				
<u> </u>	[11000 Babienel				

At page 24 please replace Table 1K with the following:

Table 1K. ClustalW Analysis of NOV1

- 1) Novel NOV1a (SEQ ID NO:2)
- 2) Novel NOV1b (SEQ ID NO:4)
- 3) Novel NOV1c (SEQ ID NO:6)
- 4) Novel NOV1d (SEQ ID NO:8)
- 4) gi|13569928|ref|NP-112217.1| gi|13569928| NP 112217 a disintegrin-like and metalloprotease with thrombospondin type 1 motif, 12 [Homo sapiens] (SEQ ID NO:30)
- 5) <u>gi|11493589|gb|AAC35563.1|AF163762_1</u> gi|11493589| <u>AAG35563</u> (AF163762) zinc metalloendopeptidase [Homo sapiens] (SEQ ID NO:31)
- 6) gi|10645199|ref|NP 055087.1| gi|10645199| NP 055087 a disintegrin and metalloprotease with thrombospondin motifs-7 preproprotein; a disintegrin-like and metalloprotease (reprolysin type) with thrombospondin type 1 motif, 7 [Homo sapiens] (SEQ ID NO:32)
- 7) gi|15309931|ref|XP 054419.1| gi|15309931| XP 054419 a disintegrin-like and metalloprotease domain with thrombospondin type I repeats 10 [Homo sapiens] (SEQ ID NO:33)
- 8) gi | 7656869 | ref | NP 055088.1 | gi | 7656869 | NP 055088 a disintegrin and metalloprotease with thrombospondin motifs-6 preproprotein; a disintegrin-like and metalloprotease (reprolysin type) with thrombospondin type 1 motif, 6 [Homo sapiens] (SEQ ID NO:34)

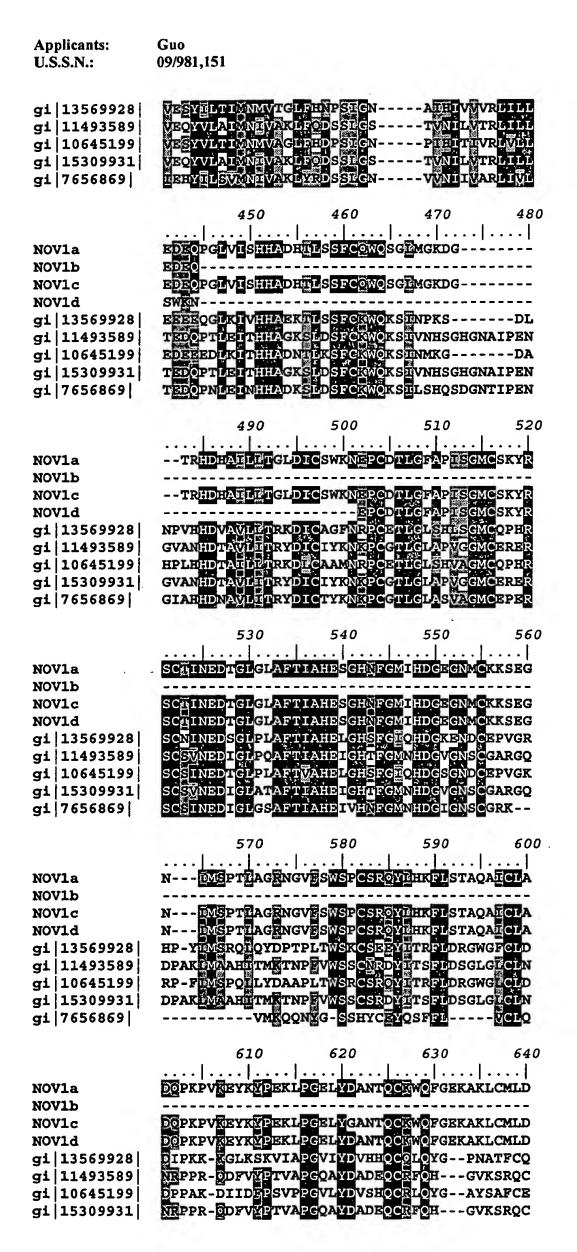
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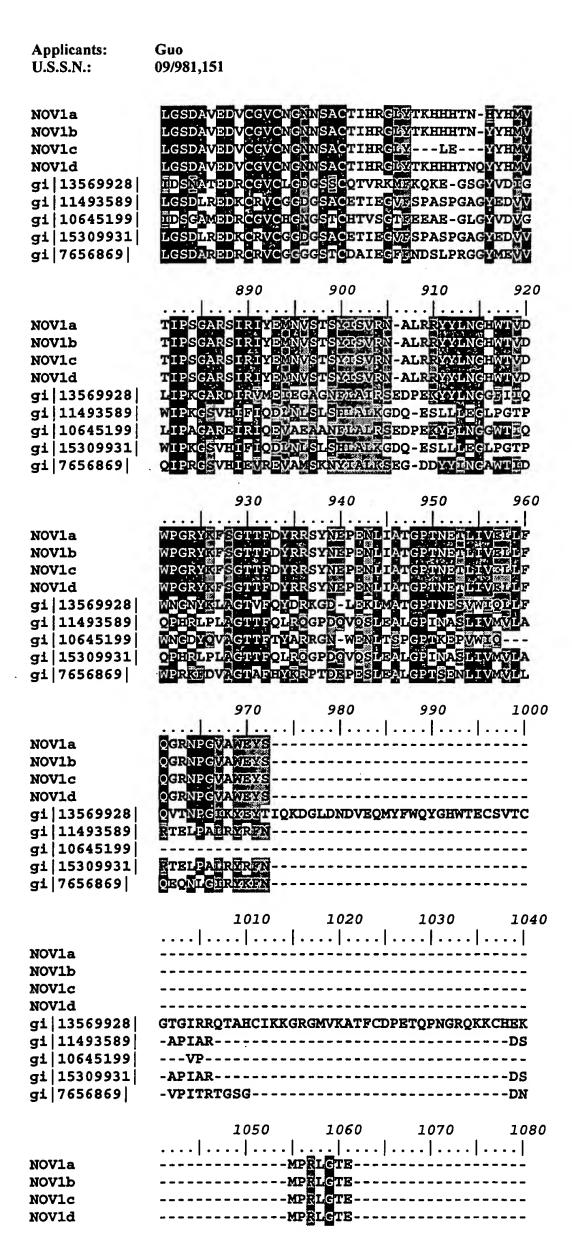
09/981,151

	10	20	30	40
-		.		
NOV1a	MKPRARGWRGLAALW			
NOV1b	MKPRARGWRGLAALW	WITTYÖNYRÖNS	PGKSHQKGN	RGSGQ
NOV1c NOV1d	MEDDADOWDOT ARTM	MT T 301/37/01/0	DCDCVODCX	 pagan
gi 13569928	MKPRARGWRGLAALW	MILING AND CAS	PGRSAQRGN.	KG2GQ
gi 13303528 gi 11493589				
gi 10645199				
gi 15309931				
gi 7656869				
- ,				
	50	60	70	80
		.		
NOV1a	LEASPPRLLSRGPRR			
NOV1b	LEASPPRLLSRGPRR			
NOV1c		-AAAAP		GSPSV
NOV1d	LEASPPRLLSRGPRR		- +	
gi 13569928 gi 11493589	MPCAQKSWIIAND	2 A AMÖTTIML GW	TC1GKQFQF(JF VRF
gi 10645199	MPGGPSPRSPAP	LLRPLLLLLCA	LAPG-APGP	APGRA
gi 15309931				
gi 7656869	ME	ILWKTLTWILS	LIMASSEFH:	SDHRL
• •				
	90	100	110	120
		.		
NOV1a	PERPASSSTRGAAGLI			
NOV1b	PERPASSSTRGAAGLI	DGKGRDMDEAG	NHRSQQTNT	GTENQ
NOV1c NOV1d	PRPPP PERPASSSTRGAAGLI	nakabu m ubya		 2TENO
gi 13569928	PDRRQEHFIKGL	DAAGRDMDAAG	MHKSQQINIC	
gi 11493589	RSQDEFLSSL			
gi 10645199	TEGR			
gi 15309931				
gi 7656869	SYSSQEEFLTYL			
	130	140	150	160
				350
NOV1a NOV1b	TLHVLT-OYDEVSAY	1000		
NOV16	TLHVLTREYDEVSAY	SVDHRGDVVSE	EMHHOR - R	AVA
NOV1d	TLHVLT-QYDEVSAY	SVDHRGDAVSH	EMHHOR - R	TAVA
gi 13569928	PEYHVVGPVI	200 200 200 200		
gi 11493589	esyetafpti			
gi 10645199	AALDIIVHPVI	RVDAGGS FFSY	elwpr-ai	RKRD
gi 15309931				
gi 7656869	EHYQETIPI	rvdongadisf	TUKND-KHS	RRRS
	170	180	190	200
NOV1a	fid ever Put	KGPRHDFHM	nii preenin	DCET
NOV1b		KGPRHDFHM		
NOV1C		CRARELRLC		
NOV1d	VSEVES-DHL			
gi 13569928	DGSEDW-VYY	AND THE RESERVE OF THE PERSON	500 Table 10 10 10 10 10 10 10 10 10 10 10 10 10	388
gi 11493589	ATAESRIFY	ASPSTHFLL	NLTRSSRLL	GHVS
gi 10645199	VSVRRDAPAFYI	QYRGRELRF	NLTAÑOHLI.	PGFV
gi 15309931				
gi 7656869	MDPIDPQQAVSKIFF	KI SAYGKHFHL	NITLATORUS	KHET
	010	222	224	240
	210	220 	230	240
	• • • • • • • • •	.	• • • • • • • •	1

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NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	VOTLGATETKSVQTLPP-EDFCFYQGSLRSHRNSPSHG VOTLGATETKSVQTLPP-EDFCFYQGSLRSHRNSPSHG NLSNVRSHWREQHASKRII-TNAMLGESALASTRKSN VOTLGATETKSVQTLPP-EDFCFYQGSLRSHRNSPSHG MEKR-YGNLSHVKMMASSAPLCHLSGTVLQQGTRVGTA VEYWTREGLAWQRAARPHCLYAGHLQGQASSS-HV SETRRAGGLGRAHIRAHTPACHLLGEVODPELEGGLA VEYWGXDGPQWKHDFLDNCHYTGYLODQRSTT-KV
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	250 260 270 280 GKFCEGSTRTLKLCNSQKCPRDSVDFRAAQCAEHNSRRFR GKFCEGSTRTLKLCNSQKCPRDSVDFRAAQCAEHNSRRFR CVFFL-SFYFFQSGMIRTEEADYFLRPLPSHLSWKLG GKFCEGSTRTLKLCNSQKCPRDSVDFRAAQCAEHNSRRFR ALSACHGLTGFFQLPHGDFFIEPVKKHPLVEG- AISTCGGLHGLIVADEEEYLIEPLHGGPKGS AISACDGLKGVFQLSNEDYFIEPLDSAPARPG- ALSNCVGLHGVIATEDEEYFIEPLKNTTEDSKH
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	290 300 310 320 GRHYKWKPYTQVEXDLCKLYCTAEGFDFFFSLSNKVKDGT GRHYKWKPYTQVEQDLCKLYCTAEGFDFFFSLSNKVKDGT RAAQGSSPSHVLYKREVLVTSRTWELAHQPLHSSDLRLGL GRHYKWKPYTQVEADLCKLYCTAEGFDFFFSLSNKVKDGTGYHPHIVYRRQKVPETKEP-TCGTKDSV RSPEESGPHVVYKRSSLRHPHTDTACGV-RDEKPWKGRPWHAQPHVVYKRQAPERTAQRGDSSAPSTCGVQVYP FSYENGHPHVIYKKSALQQRHTYDHSHCGVSDFTRSGKPW
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	330 340 350 360
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	370 380 390 400 YKSCLRHKRSILKSHRNEELNVETLVVVDKKMMONHGHEN YKSCLRHKRSILKSHRNEELNVETLVVVDKKMMONHGHEN YKSCLRHKRSILKSHRNEELNVETLVVVDKKMMONHGHEN YKSCLRHKRSILKSHRNEELNVETLVVVDKKMMONHGHEN WERHNLPSKSISKERWETLVVADTKMEEYHGSEN LGNETERGOPGLKRSVSKERWVETLVVADKMMVAYHGRRD OÖWRRPRLRRLHORSVSKEKWVETLVVADAKMWEYHGQPQ
NOV1a NOV1b NOV1c NOV1d	410 420 430 440



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gi 7656869	sRxнно
	650 660 670 680
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	FKKDICKALWCHRIGRKCETKFYPAAEGTECGHDMWDICKALWCHRIGRKCETKFYPAAEGTICGHDMW FKKDICKALWCHRIGRKCETKFYPAAEGTICGHEHG FKKDICKALWCHRIGRKCETKFYPAAEGTICGHEHG FKKDICKALWCHRIGRKCETKFYPAAEGTICGHDMW EVENYCOTLWCSVKG-FCRSKLDAAASGTOCGEKKW KYGSYGSELWCLSKSNRCITNSSPAAEGTFCOTHTIDKGW DMDNYCHTLWCSVGT-TCHSKLDAAVSGTRCGENKW KYGSYCSELWCLSKSNRCITNSSPAAEGTFCOTHTIDKGW LFRSYCKELWCLSKSNRCVTNSSPAAEGTFCOTGNIEKGW
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	CRGGCCVKYGDEGPKPTHGHWSDWSSWSPCSRTCGGGVSHCRGGCVKYGDEGPKPTHGHWSDWSSWSPCSRTCGGGVSHCRGGCVKYGDEGPKPTHGHWSDWSSWSPCSRTCGGGVSHCRGGCVKYGDEGPKPTHGHWSDWSSWSPCSRTCGGGVSHCRGGCVKYGDEGPKPTHGHWSDWSSWSPCSRTCGGGVSHCRGGCVKYGDEGPKPTHGHWSDWSSWSPCSRTCGGGVSHCMAGKCTVCKK-PESIPGGWGRWSPWSHCSRTCGAGVSSCYKRVCVPTGSR-PEGVDGAWGPWTPWGDCSRTCGGGVSSCLSGECVPVGFR-PEAVDGGWSGWSAWSICSRSCGMGVQSCYKRVCVPTGSR-PEGVDGAWGPWTPWGDCSRTCGGGVSSCYKRVCVPTGSR-PEGVDGAWGPWTPWGDCSRTCGGGVSSCYQGDCVPTGTW-POSIDGGWGPWSLWGECSRTCGGGVSS
	730 740 750 760
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	RSR-LCTNPRPSHGGKECEGSTRTLELCNSOKCPRDSVDF RSR-LCTNPRPSHGGKECEGSTRTLELCNSOKCPRDSVDF RSRSQNTHSRPSHGGKECEGSTRTLELCNSOKCPRDSVDF RSR-LCTNPRPSHGGKECEGSTRTLELCNSOKCPRDSVDF RSR-LCTNPRPSHGGKECEGSTRTLECNSOKCPRDSVDF AER-LCNNPEPKFGGKECTGERERYRLCNVHPCRSENPTF SSR-HCDSPRPTILGGKECLGERERHESCNTDDCPPGSODF AER-QCTOPTPKYKGEECVGERERFRLCNLOACPAGRPSF SSR-HCDSPRPTILGGKECLGERERHESCNTDDCPPGSODF SLR-HCDSPAPSGGGKECLGERERYESCNTDDCPPGSODF SLR-HCDSPAPSGGGKECLGERERYESCNTDDCPPGSODF
	770 780 790 800
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	RAAQCAEHISRRFRGRHYKWKPQDLCKLYCHAEG RAAQCAEHISRRFRGRHYKWKPQDLCKLYCHAEG RAAQCAEHISRRFRGRHYKWKP
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	### ##################################
	850 860 870 880



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gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	ACPPRWWAGEWEACSATC PHGEKKRTVLCIQTMVSDEQA LPPYSWHYAPWTKCSA CAGGSQV-QAVECRNQLDSSAGG LPPYSWHYAPWTKCSA CAGGSQV-QAVECRNQLDSSA EVGFTWNHQPWSECSATCAGG
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	1090 1100 1110 1120
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	1130 1140 1150 1160
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	1170 1180 1190 1200 RRVLKPNKGTISNGKNPPTLKPVPPPTSRPRMLTTPTGPERVSAAEEKALDDSACPQ
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	1210 1220 1230 1240
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931	1250 1260 1270 1280

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	<u> </u>
gi 7656869	KET <mark>ill</mark>
	1290 1300 1310 1320
NOV1a	
NOV1b	
NOV1c	
NOV1d	GLVATTTSGSGLSSSRNPITWPVTPFYNTLTKGPEMEIHS
gi 13569928 gi 11493589	GLVAIIISGSGDSSSKAPIIWFVIFFINIDIKGPEMEINS
gi 10645199	GIHR
gi 15309931	PAT
gi 7656869	· · ·
91/7030003/	
	1330 1340 1350 1360
NOV1a	
NOV1b	
NOV1c	
NOV1d	
gi 13569928	GSGEEREQPEDKDESNPVIWTKIRVPGNDAPVESTEMPLA
gi 11493589	MRCNLRRCPP
gi 10645199	EAGGHDEVPP
gi 15309931	MRCNLRRCPP
gi 7656869	
	1370 1380 1390 1400
170774 -	
NOV1a NOV1b	
NOV16	
NOV1d	
qi 13569928	PPLTPDLSRESWWPPFSTVMEGLLPSQRPTTSETGTPRVE
gi 11493589	
gi 10645199	
gi 15309931	
91 122003331	
gi 7656869	
	1410 1420 1430 1440
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gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928	
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gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931	
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gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931	GMVTEKPANTLLPLGGDHQPEPSGKTANRNHLKLPNNMNQ
gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931	GMVTEKPANTLLPLGGDHQPEPSGKTANRNHLKLPNNMNQ 1450 1460 1470 1480
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gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	GMVTEKPANTLLPLGGDHQPEPSGKTANRNHLKLPNNMNQ 1450 1460 1470 1480
gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 NOV1a NOV1b NOV1c NOV1d	GMVTEKPANTLLPLGGDHQPEPSGKTANRNHLKLPNNMNQ 1450 1460 1470 1480
gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928	GMVTEKPANTLLPLGGDHQPEPSGKTANRNHLKLPNNMNQ 1450 1460 1470 1480
gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589	GMVTEKPANTLLPLGGDHQPEPSGKTANRNHLKLPNNMNQ 1450 1460 1470 1480
gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 7656869 NOV1a NOV1b NOV1c NOV1c NOV1d gi 13569928 gi 11493589 gi 11645199	GMVTEKPANTLLPLGGDHQPEPSGKTANRNHLKLPNNMNQ 1450 1460 1470 1480 TKSSEPVLTEEDATSLITEGFLLNASNYKQLTNGHGSAHW
gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 11493589 gi 11645199 gi 15309931	GMVTEKPANTLLPLGGDHQPEPSGKTANRNHLKLPNNMNQ 1450 1460 1470 1480 TKSSEPVLTEEDATSLITEGFLLNASNYKQLTNGHGSAHW
gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 7656869 NOV1a NOV1b NOV1c NOV1c NOV1d gi 13569928 gi 11493589 gi 11645199	GMVTEKPANTLLPLGGDHQPEPSGKTANRNHLKLPNNMNQ 1450 1460 1470 1480 TKSSEPVLTEEDATSLITEGFLLNASNYKQLTNGHGSAHW
gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 11493589 gi 11645199 gi 15309931	GMVTEKPANTLLPLGGDHQPEPSGKTANRNHLKLPNNMNQ 1450 1460 1470 1480 TKSSEPVLTEEDATSLITEGFLLNASNYKQLTNGHGSAHW
gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 7656869 NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 11493589 gi 11645199 gi 15309931	GMVTEKPANTLLPLGGDHQPEPSGKTANRNHLKLPNNMNQ 1450 1460 1470 1480 TKSSEPVLTEEDATSLITEGFLLNASNYKQLTNGHGSAHW

Applicants:

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Applicants: U.S.S.N.:	Guo 09/981,151
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	IVGNWSECSTTCGLGAYWKRVECTTQMDSDCAAIQRPDPA VAGEWGECSAQCGVGQRQRSVRCTS
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	1530 1540 1550 1560 KRCHLRPCAGWKVGNWSKCSRNCSGGFKIREIQCVDSRDH
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	1570 1580 1590 1600 RNLRPFHCQFLAGIPPPLSMSCNPEPCEAWQVEPWSQCSR
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	1610 1620 1630 1640 SCGGGVQERGVFCPGGLCDWTKRPTSTMSCNEHLCCHWAT KCDSPTPG HCWAT KCDSPTPG
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	1650 1660 1670 1680 GNWDLCSTSCGGGFQKRIVQCVPSEGNKTEDQDQCLCDHK DGPEECKDVN
NOVla NOVlb NOVlc NOVld	1690 1700 1710 1720

Applicants:	Guo	
U.S.S.N.:	09/981,151	
gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869	PRPPEFKKCNQQACKKSADLLCTKDKLSASFCQTLKAN	?QF IAL
NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931	1730 1740 1750	(SEQ ID NO:2) (SEQ ID NO:4) (SEQ ID NO:6) (SEQ ID NO:8) (SEQ ID NO:30) (SEQ ID NO:31) (SEQ ID NO:32) (SEQ ID NO:33)
gi 7656869		(SEQ ID NO:34)

At page 32, please replace paragraph at lines 31-42 with the following new paragraph:

The presence of identifiable domains in NOV1, as well as all other NOVX proteins, was determined by searches using software algorithms such as PROSITE, DOMAIN, Blocks, Pfam, ProDomain, and Prints, and then determining the Interpro number by crossing the domain match (or numbers) using the European Bioinformatics Institute Interpro. website (http://www.ebi.ac.uk/interpro). DOMAIN results for NOV1 as disclosed in Tables 1L-IO, were collected from the Conserved Domain Database (CDD) with Reverse Position Specific BLAST analyses. This BLAST analysis software samples domains found in the Smart and Pfam collections. For Table 1E and all successive DOMAIN sequence alignments, fully conserved single residues are indicated by black shading or by the sign (|) and "strong" semi-conserved residues are indicated by grey shading or by the sign (+). The "strong" group of conserved amino acid residues may be any one of the following groups of amino acids: STA, NEQK (SEQ ID NO: 161), NHQK (SEQ ID NO: 162), NDEQ (SEQ ID NO: 163), QHRK (SEQ ID NO: 164), MILV (SEQ ID NO: 165), MILF (SEQ ID NO: 166), HY, FYW.

At page 33, please replace Table 1L with the following:

Table 1L. Domain Analysis of NOV1

gnl|Pfam|pfam01421, Reprolysin, Reprolysin (M12B) family zinc metalloprotease. The members of this family are enzymes that cleave peptides. These proteases require zinc for catalysis. Members of this family are also known as adamalysins. Most members of this family are snake venom endopeptidases, but there are also some mammalian proteins, and fertilin. Fertilin and closely related proteins appear to not have some active site residues and may not be active enzymes. (SEQ ID NO:76)

CD-Length = 199 residues, 95.0% aligned Score = 122 bits (307), Expect = 7e-29

```
DKKMMQNHG--HENITTYVLTILNMVSALFKDGTIGGNINIAIVGLILLEDEQPGLVISH
Query:
       378
                   +|
                          | |+|+|+ +++
                                                 || + +||| + |
            DHGMFTKYGSDLNKIRQRVHQIVNLVNEIYRPL----NIRVVLVGLEIWSDGDK-ITVQG
Sbjct:
       10
            {\tt HADHTLSSFCQWQSGLMGKDGTRHDHAILLTGLDICSWKNEPCDTLGFAPISGMCSKYRS}
Query:
       436
             |+ || |+|+
                           + |
                                   ||+| ||||+|
                                                       +|+| | + ||||
            DANDTLHRFLEWRETDLLKR-KSHDNAQLLTGIDF-----DGNTIGAAYVGGMCSPKRS
Sbjct:
       65
                                                                          117
            CTINED---TGLGLAFTIAHESGHNFGMIHDGEGN-MCKKSEGNIMSPTLAGRNGVFSWS
                                                                          551
Query:
       496
                       | +| |+||| ||| ||
              + +|
                                                 +
                                                      | ||+| +
            VGVVQDHSPIVLLVAVTMAHELGHNLGMTHDDINKCTCEGGGGCIMNPVASSSPGKK-FS
Sbjct:
       118
            PCSRQYLHKFLSTAQAICLADQ 573
Query:
                                         (aa 378-573 of SEQ ID NO:2)
                    +||+ + || ++
             Ш
       177 NCSMDDYQQFLTKGKPQCLLNK 198
Sbjct:
                                         (SEQ ID NO:76)
```

At page 33, please replace Table 1M with the following:

Table 1M. Domain Analysis of NOV1 gnl|Smart|smart00209, TSP1, Thrombospondin type 1 repeats; Type 1 repeats in thrombospondin-1 bind and activate TGF-beta. (SEQ ID NO:77) CD-Length = 51 residues, 100.0% aligned Score = 63.5 bits (153), Expect = 5e-11

At page 33, please replace Table 1N with the following:

Table 1N. Domain Analysis of NOV1

gnl|Pfam|pfam00090, tsp_1, Thrombospondin type 1 domain. (SEQ ID NO:78)
CD-Length = 48 residues, 100.0% aligned
Score = 54.7 bits (130), Expect = 2e-08

Query: 669 SDWSSWSPCSRTCGGGVSHRSRLCTNPNPSHGGKFCEGSTRTLKLCNSQKC 719 (of SEQ ID NO:2)

| | | | | | | | | + | | | + + | |

Sbjct: 1 SPWSEWSPCSVTCGKGIRTRQRTCNSP---AGGKPCTGDAQETEACMMDPC 48 (SEQ ID NO:78)

At page 34, please replace Table 10 with the following:

Table 10. Domain Analysis of NOV1

gnl|Pfam|pfam01562, Pep_M12B_propep, Reprolysin family propeptide.
This region is the propeptide for members of peptidase family M12B.
The propeptide contains a sequence motif similar to the "cysteine switch" of the matrixins. This motif is found at the C terminus of the alignment but is not well aligned. repeats. (SEQ ID NO:79)
CD-Length = 117 residues, only 39.3% aligned
Score = 38.5 bits (88), Expect = 0.002

Query: 180 LRTSSSLVAPGFIVQTLGKTGTKSVQTLPPEDFCFYQGSLRSHRNS 225 (of SEQ ID NO:2)

| + | | + | | | | | | + + | | + | | + | |

Sbjct: 2 LEKNRSLLAPDFTVTTYDDDGTLVTEHPLIQDHCYYQGYVEGYPNS 47 (SEQ ID NO:79)

At page 41 please replace Table 2C with the following:

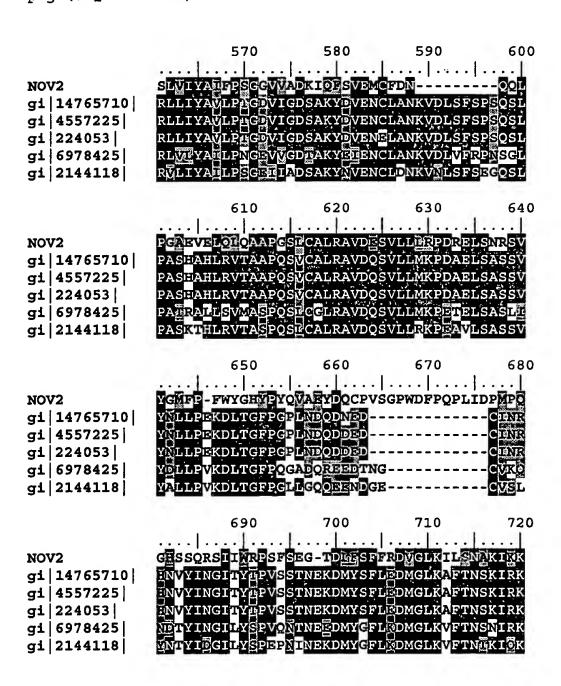
Table 2C. BLAST results for NOV2					
Gene Index/	Protein/	Length	Identity	Positives	Expect
Identifier	Organism	(aa)	(%)	(%)	
gi 14765710 ref XP- 006925.4 XP 006925	alpha 2 macroglobulin precursor [Homo sapiens]	1474	593/1486 (39%)	870/1486 (57%)	0.0
<u>gi 4557225 ref NP 0</u> 00005.1 NP 000005	alpha 2 macroglobulin precursor [Homo sapiens]	1474	591/1486 (39%)	869/1486 (57%)	0.0
gi 224053 prf 1009 174A 1009174A	macroglobulin alpha2 [Homo sapiens]	1450	585/1471 (39%)	861/1471 (57%)	0.0

gi 6978425 ref NP 0 36620.1 NP 036620	alpha-2- macroglobulin [Rattus norvegicus]	1472	578/1483 (38%)	867/1483 (57%)	0.0
gi 2144118 pir JC5 143 JC5143	alpha- macroglobulin precursor - guinea pig	1476	570/1495 (38%)	858/1495 (57%)	0.0

At page 41 please replace Table 2D with the following:

Table 2D. ClustalW Analysis of NOV2

- 1) NOV2 (SEQ ID NO:10)
- 2) <u>gi|14765710|ref|XP 006925.4|</u> <u>gi|14765710|</u> <u>XP 006925</u> alpha 2 macroglobulin precursor [Homo sapiens] (SEQ ID NO:35)
- 2) <u>gi|4557225|ref|NP 000005.1|</u> <u>gi|4557225|</u> <u>NP 000005</u> alpha 2 macroglobulin precursor [Homo sapiens] (SEQ ID NO:36)
- 3) <u>gi|224053|prf||1009174A</u> gi|224053| 1009174A macroglobulin alpha2 [Homo sapiens] (SEQ ID NO:37)
- 4) <u>gi|6978425|ref|NP 036620.1|</u> <u>gi|6978425|</u> NP 036620 alpha-2-macroglobulin [Rattus norvegicus] (SEQ ID NO:38)
- 5) gi|2144118|pir||JC5143 gi|2144118| JC5143 alpha-macroglobulin precursor guinea pig (SEQ ID NO:39)



	730	740	750	760
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	PVDCSHRSPEYST PKVCPOLOOVEVHO PKVCPOLOOVEVHO PKVCPOLOOVEWHO PKVCPOLOOVEWHO PKVCERLEDNEGI	GPEGURVGÖYES GPEGURVGÖYES GPEGURVGÖYES PAAYHLVSQSH	DVMGRGHARI DVMGRGHARI DVMGRGHARI DAFLE	VHV
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	770 LHOARDSOVRREPHTETVREPHTETVREPHTETVR SSESPTETRR AATYSEPPKETVR	YFPETWIWDL\ YFPETWIWDL\ YFPETWIWDL\ YFPETWIWDL\	/VVŠSAGVAE /VVŠSAGVAE /VVŠSAGVAE /VVŠSAGVAE	/GVIVP /GVIVP /GVIVP /EVIVP
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	810 DAITEWKAMSFCT DTITEWKAGAFCL DTITEWKAGAFCL DTITEWKAGAFCL DTITEWKAGAFCL DTITEWKAGAFCL	SEDAGEGESSTA SEDAGEGESSTA SEDAGEGESSTA SUDTGEGESPVV	SLRAFOPFF SLRAFOPFF SLRAFOPEE OFGAFOPFF	/ELTMP /ELTMP /ELTMP /ELTMP
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	850 YSVVRGESFRLMA YSVIRGEAFTLKA YSVIRGEAFTLKA YSVIRGEAFTLKA YSVIRGEAFTLKA YSVIRGEAFTLKA	IVLNYLP <mark>K</mark> CIRV IVLNYLPKCIRV IVLNYLPTCIRV	/SVOLEASPAI /SVOLEASPAI /SVOLEASPAI /SVOLEASPOI	LAVPV LAVPV LAVPV LAAPE
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	890 ADSOTSSCICADO EKEOAPHCICANG EKEOAPHCICANG EKEOAPHCICANG EKEOAPHCICANG EKEORSHCICANO AKEOESICÜCGNE	RQTVSWAVTPKS RQTVSWAVTPKS RQTVSWAVTPKS RHT <mark>A</mark> SWAV <mark>T</mark> PKS	LĠŊŲŊŦŢŲĠĮ LĠŊŲŊŦŢŲĠĮ LĠŊŲŊŦŢŲĠĮ LĠŊŲŊŦŢŲĠĮ	leales Leales Leales Leal <mark>n</mark> s
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	930 MEPCGGOKGFVPOT ELCGTEVPSVPE ELCGTEVPSVPE ELCGTEVPSVPE ELCGNEVPVVPE SELCGNEKTVVPT	GRKDTÖIKPLI GRKDTÖIKPLI GRKDTÖIKPLI GKKDTÖIKS	VEPEĞLEKET VEPEĞLEKET VEPEĞLEKET VEPEĞLE <mark>N</mark> EV	TFNSL TFNSL TFNSL TFNSL
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	970 LCPKGGKVASES LCPSGGEVS-EEL LCPSGGEVS-EEL LCPMGAEVS-EL RVSDTTVS-EKL	SLKLP <mark>P</mark> NVVEES SLKLP <mark>P</mark> NVVEES SLKLP <mark>S</mark> EVVEES	ĀRASŲSŲLGI ARASŲSŲLGI ARASŲSŲLGI ARASV Ū VLGI	ILGSĀ ILGSĀ ILGSĀ ILGSĀ
	1010	1020	1030	1040



Applicants: U.S.S.N.:	Guo 09/981,151
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	MONTONLLOMPYGCGEONMVLFAPITYVLOYLEGAGLITE MONTONLLOMPYGCGEONMVLFAPNIYVLDYLNETOOLTP MONTONLLOMPYGCGEONMVLFAPNIYVLDYLNETOOLTP MONTONLLOMPYGCGEZNMVLFAPNIYVLDYLNETOOLTP MONTOBLIZMPYGCGEONMVLFAPNIYVLDYLNETOOLTQ MONTOBLIZMPYGCGEONMVLFAPNIYVLDYLNETOOLTQ MONTOBLIZMPYGCGEONMVLFAPNIYVLDYLNETOOLTP
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	1050 1060 1070 1080 EIRSRAUGELEIGYONZLMYKHSIGSYSAFGERDGNGN EIKSKAIGYLNTGYOROLNYKHYDGSYSTFGERYGRNOGN EIKSKAIGYLNTGYOROLNYKHYDGSYSTFGERYGRNOGN EIKSKAIGYLNTGYOROLNYKHYDGSYSTFGERYGRNOGN EIKKAIAYLNTGYOROLNYKHYDGSYSTFGERYGRNOGN EIKKKAIAYLNTGYOROLNYKHRDGSYSAFGDEPGRNHAN EIKSKAI
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	1090 1100 1110 1120
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	1130 1140 1150 1160 CVANVGNLL TAKKGGVDDEVSL AYVTAALLENGKDVDD CFRSSGSLLNNAIKGGVEDEVTLSAYITIALLE PLIVTH CFRSSGSLLNNAIKGGVEDEVTLSAYIKIALLE PLIVTH CFRSSGSLLNNAIKGGVEDEVTLSAYIKIALLE SLPVTH CFRSSGSLLNNAIKGGVEDE SLSAYITIALLE SLPVTH CFWSSGSLLNNAIKGGVEDE SLSAYITIALLE SLPVTH
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	1170 1180 1190 1200 PVVSQGLRCLKNSATST
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	1210 1220 1230 1240 EMDIRNITLKOLDOOAIISGESIYWSOKPTPSSNASPYSE NODERKEYLKSLNEEAVKEDNSVHWERPOKPKAPVGHIYE NODERKEYLKSLNEEAVKEDNSVHWERPOKPKAPVGHIYE NODERKEYLKSLNEEAVKEDNSVHWERPOKPKAPVGHIYE NODERKEYLKSLNEEAVKEDNSVHWERPOKPKAPVGHIYE NODERKEYLKSLNEEAVKEDISSVHWERPOKPKVSVGLWYO NOOTEKEILKSLEEEGVKEDNSVHWERPOKPKVSEGFLOK
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	1250 1260 1270 1280 PAAVDVEETAYALLAOLT- PELTO CERAKAT SIVAWI POAPSAEVEMTSYVLLAYLTAO PAPTSEDLTSAT NIVKWE POAPSAEVEMTSYVLLAYLTAO PAPTSEDLTSAT NIVKWE POAPSAEVEMTSYVLLAYLTAO PAPTSEDLTSAT NIVKWE POATSAEVEMTSYVLLAYLTA PAPTO EDLTSAT NIVKWE SOAPSAEVEMTSYVLLAYLTA PAPTPEDLTSAT DIVNWA 1290 1300 1310 1320







Applicants: U.S.S.N.:	Guo 09/981,151
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	AKQENAYGGFSSTQDTVVALQALKKYATTAYMPS-EEINE TKQQNAQGGFSSTQDTVVALHALSKYGAATFTRTGKAAQV TKQQNAQGGFSSTQDTVVALHALSKYGAATFTRTGKAAQV TKQQNAQGGFSSTQDKVVALHALSKYGAATFTRTGKAAQV TKQQNSHGGFSSTQDTVVALHALSKYGSÄTFTRAKKAAQV TKQQNSHGGÄSSTQDTVVALHALSKYAAATFTRTEKAAQV
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	1330 1340 1350 1360 VVASEENFORTFNEOSVNRLVFOODELPNVPGMYHTEASG TI SSGTFSSKFOV NNNRLLLOOVSLPEPPGSYSVKVTG TI SSGTFSSKFOV NNNRLLLOOVSLPEPPGSYSVKVTG TI SSGTFSSKFOV NNNRLLLOOVSLPEPPGSYSVKVTG TI SSGTFSSKFOV NNNGLLLOOVSLPEPPGSYSVKVTG TI SSGTFSSKFOV NNNGLLLOOVSLPEPPGSYSVKVTG TI SSGTFSSKFOV NNNGLLLOOVSLPEPPGSYSVKVTG TI SSGTFSSKFOV NNNKLLLOOVSLPTVPGSYSVKVTG
	1370 1380 1390 1400
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	©GCVYVOTVLEYNILPETNMKTFELSVEIGKARCEOPTEP EGCVYLOTSLKYNILPEKEEEPFALGVOTLPOTCDEPKAH EGCVYLOTSLKYNILPEKEEFPFALGVOTLPOTCDEPKAH EGCVYLOTSLKYNILPEKEEFPFALGVOTLPOTCEEPKAH EGCVYLOTSLKYSVLPREEEFPFAUVVOTLPGTCEEPKAH EGNVYLOTSLKYNVPSEKGNFPFALEAETVEOACDGPKAH
	1410 1420 1430 1440
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	RSLT FEBRUSY GSRSSSNMAIV VKM SGFSPEGTNOF TSFQISLSVSYTGSRS SNMAIVDVKMVSGFIPLKPTVKM TSFQISLSVSYTGSRS SNMAIVDVKMVSGFIPLKPTVKM TSFQISLSVSYTGSRS SNMAIVDVKMVSGFIPLKPTVKM TSFQISLSUSYTGSRS SNMAIADVKMVSGFIPLKPTVKM TSFQISLSVSYTGSRS SNMAIVDVKMVSGFIPLKPTVKM
	1450 1460 1470 1480
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	LLOPLVK VEFGUET IN IYLD BUKNTOUYTETISOSVL LERSNHVSRTEVS NHVLIYLDKVSNOTUSL FFTVRÖDVP LERSNHVSRTEVS NHVLIYLDKVSNOTUSL FFTVRÖDVP LERSNHVSRTEVS NHVLIYLDKVSNOTUSL FFTVRÖDVP LERSVHVSRTEVS NHVLIYLDKVSNOTUSL SFTVOODUP LERSVHVSRTEVS NHVLIYLDKVSNOTUSL SFTVOODUS
	1490 1500 1510 1520
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	VENLKPATEKVYDYYLPGSFKESOYTIVWSMINDSIVDSV VRDLKPAIVKVYDYYETGEFAIAEYNAPCSKOLGNA VRDLKPAIVKVYDYYETDEFAIAEYNAPCSKOLGNA URDLKPAIVKVYDYYETDEFAIAEYNAPCSKOLGNA URDLKPANVKVYDYYEKOEFANANYSAPCSKOLGNA URDLKPANVKVYDYYEKOEFANANYSAPCSKOPGNA VRDLKPAIEKVYDYYETNEFAIAEYNAPCSKOPGNA
NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118	1530 1540 ARHPEPPPFKTEAFIPSLPGSVNN (SEQ ID NO:10) (SEQ ID NO:35) (SEQ ID NO:36) (SEQ ID NO:37) (SEQ ID NO:38) (SEQ ID NO:39)



Pages 45, please replace Table 2E with the following

Table 2E Domain Analysis of NOV2

gnl|Pfam|pfam00207, A2M, Alpha-2-macroglobulin family. This family
includes the C-terminal region of the alpha-2-macroglobulin family.
(SEQ ID NO:80)
CD-Length = 751 residues, 98.5% aligned
Score = 563 bits (1451), Expect = 2e-161

Query:	728	EDSQVRQYFPETWLWDLFPIGNSGKEAVHVTVPDAITEWKAMSFCTSQSRGFGLSPTVGL + + + + + + + + + + +	787
Sbjct:	4	DDITIRSYFPESWLWEVEEVDRSPVLTVNITLPDSITTWEILAVSLSNTKGLCVADPVEL	63
Query:	788	TAFKPFFVDLTLPYSVVRGESFRLTATIFNYL-KDCIRVQTDLAKSHEYQLESWADSQTS	846
Sbjct:	64	TVFQDFFLELRLPYSVVRGEQVELRAVLYNYLPSQDIKVVVQLEVEPLCQAG	115
Query:	847	SCLCADDAKTHHWNITAVKLGHINFTISTKILDSNEPCGGQKGFVPQKGRSDTLIKPVLV	906
Sbjct:	116	FCSLATQRTRSSQSVRPKSLSSVSFPVVVVPLASGLSLVEVVASVPEFFVKDAVVKTLKV	175
Query:	907	KPEGVLVEKTHSSLLCPKGGKVASESVSLELPVDIVPD-STKAYVTVLGDIMGTALQ +	962
Sbjct:	176	EPEGARKEETVSSLLLPPEHLGGGLEVSEVPALKLPDDVPDTEAEAVISVQGDPVAQAIQ	235
Query:	963	NLDGLVQMPSGCGEQNMVLFAPIIYVLQYLEKAGLLTEEIRSRAVGFLEIG	1013
Sbjct:	236	NTLSGEGLNNLLRLPSGCGEQNMIYMAPTVYVLHYLDETWQWEKPGTKKKQKAIDLINKG	295
Query:	1014	YQKELMYKHSNGSYSAFGERDGNGNTWLTAFVTKCFGQAQKFIFIDPKNIQDALKW-MAG	1072
Sbjct:	296	YQRQLNYRKADGSYAAFLHRASSTWLTAFVLKVFSQARNYVFIDEEHICGAVKWLILN	353
Query:	1073	NQLPSGCYANVGNLLHTAMKGGVDDEVSLTAYVTAALLEMGKDVDDPMVSQGLRCL	1128
Sbjct:	354	QQKDDGVFRESGPVIHNEMKGGVGDDAEVEVTLTAFITIALLEAKLVCISPVVANALSIL	413
Query:	1129	KNSATSTTNLYTQALLAYIFSLAGEMDIRNILLKQLDQQAIISGESIYWSQK	1180
Sbjct:	414	KASDYLLENYANGORVYTLALTAYALALAGVLHKLKEILKSLKEELYKALVKGHWERPOK	473
Query:	1181	PTPSSNASPWSEPAAVDVELTAYALLAQLTKPSLTQKEIAKATSIVAWLAKQHNAYGGFS + + + + + + +	1240
Sbjct:	474	PKDAPGHPYSPQPQAAAVEMTSYALLALLTLLPFPKVEMAPKVVKWLTEQQYYGGGFG	531
Query:	1241	STQDTVVALQALAKYATTAYMPSE-EINLVVKSTEN-FQRTFNIQSVNRLVFQQDTLP-N	1297
Sbjct:	532	STQDTVMALQALSKYGIATPTHKEKNLSVTIQSPSGSFKSHFQILNNNAFLLRPVELPLN	591
Query:	1298	VPGMYTLEASGQGCVYVQTVLRYNILPPTNMKTFSLSVEIGKARCEQPTSPR-SLTLTIH	1356
Sbjct:	592	EGFTVTAKVTGQGTLTLVTTYRYKVLDKKNTFCFDLKIETVPDTCVEPKGAKNSDYLSIC	651
Query:	1357	TSYVGSRSSSNMAIVEVKMLSGFSPMEGTNQLLLQQPLVKKVEFGTDTLNIYLDELIK	1414
Sbjct:	652	TRYAGSRSDSGMAIADISMLTGFIPLKPDLKKLENGVDRYVSKYEIDGNHVLLYLDKVSH	711
Query:	1415	-NTQTYTFTISQSVLVTNLKPATIKVYDYYLP 1445 (of SEQ ID NO:10) +	

Sbjct: 712 SETECVGFKIHQDFEVGLLQPASVKVYDYYEP 743 (SEQ ID NO:80)

Pages 46, please replace Table 2F with the following:

Table 2F Domain Analysis of NOV2

gnl|Pfam|pfam01835, A2M_N, Alpha-2-macroglobulin family N-terminal
region. This family includes the N-terminal region of the alpha-2macroglobulin family. (SEQ ID NO:81)
CD-Length = 620 residues, 98.4% aligned
Score = 236 bits (603), Expect = 5e-63

Query:	5	LLLGMLALSPAIAEELPNYLVTLPARLNFPSVQKVCLDLSPGYSDVKFTVTLETKDKT	62
Sbjct:	2	LLWLLLLLLFFDSSLQKPRYMVIVPSILRTETPEKVCVQLHDLNETVTVTVSLHSFPGK	61
Query:	63	QKLLEYSGLKKRHLHCISFLVPPPAGGTEEVATIRVSGVGNNISFEEKKKVLIQ +	116
Sbjct:	62	RNLSSLFTVLLSSKDLFHCVSFTVPQPGLFKSSKGEESFVVVQVKGPTHTFKEKVTVLVS	121
Query:	117	RQGNGTFVQTDKPLYTPGQQVYFRIVTMDSNFVPVNDKYSMVELQDPNSNRIAQWLEVVP + + + + ++ + + + + +	176
Sbjct:	122	SRRGLVFIQTDKPIYTPGQTVRYRVFSVDENLRPLNELI-LVYIEDPEGNRVDQWEVNKL	180
Query:	177	EQGIVDLSFQLAPEAMLGTYTVAVAEGKTFGTFSVEEYVLSPFLLLLSSVLPKFK	231
Sbjct:	181	EGGIFQLSFPIPSEPIQGTWKIVARYESGPESNYTHYFEVKEYVLPSFEVS	231
Query:	232	VEVVEPKELSTVQESFLVKICCRYTYGKPMLGAVQVSVCQKANTYWYREVEREQLPDKCR + +	291
Sbjct:	232	ITPPKPFIYYDNFKEFEVTICARYTYGKPVPGVAYVRFGVKDEDGKKELLAGLE	285
Query:	292	NLSGQTDKTGCFSAPVDMATFDLIGYAY-SHQINIVATVVEEGTGVEANA-TQNIYIS +	347
Sbjct:	286	ERAKLLDGNGEICLSQEVLLKELQLKNEDLEGKSLYVAVAVIESEGGDMEEAELGGIKIV	345
Query:	348	PQMGSMTFEDTSNFYHPNFPFSGKMLLKFPQGGVLPCKNHLVFLVIYGTNGTFNQTLVTD +	407
Sbjct:	346	RSPYKLKFVKTPSHFKPGIPFFLKVLVVDPDGSPAPNVPVKVSAQDASYYSNGTTD	401
Query:	408	NNGLAPFTLETSGWNGTDVSLEGKFQMEDLVYNPEQVPRYYQNAYLHLRPFYSTTRSFLG + + + + + + + + + + + +	467
Sbjct:	402	EDGLAQFSINTSGISSLSITVRTNHKELPEEVQAHAEAQATAYSTVSLSKSYIHLS	457
Query:	468	<pre>IHRLNGPLKCGQPQEVLVDYYIDPADASPDQEISFSYYLIGKGSLVMEGQKHLNSKKKGL </pre>	527
Sbjct:	458	IERTLPCGPGVGEQANFILRGKSLGELKILHFYYLIMSKGKIVKTGREPREPG	510
Query:	528	KASFSLSLTFTSRLAPDPSLVIYAIFPSGGVVADKIQFSVEMCFDNQQL + + + + +	576
Sbjct:	511	+ + + + +	570
Query:	577	PGAEVELQLQAAPGSLCALRAVDESVLLLRPDRELSNRSVY 617 (of SEQ ID NO	:10)
Sbjct:	571	+ + +++	<u> </u>

Applicants: U.S.S.N.:

Guo 09/981,151

At page 52 please replace Table 3E with the following:

Table 3E. BLAST results for NOV3						
Gene Index/ Identifier	Protein/ Organism	Length (aa)	Identity (%)	Positives (%)	Expect	
gi 12858115 dbj BAB 31203.1 (AK018423) BAB31203	putative [Mus musculus]	373	187/310 (60%)	226/310 (72%)	6e-87	
gi 3024224 sp Q2872 7 NTCI RABIT NP 058918	ILEAL SODIUM/BILE ACID COTRANSPORTER (ILEAL NA(+)/BILE ACID COTRANSPORTER) (NA+ DEPENDENT ILEAL BILE ACID TRANSPORTER) (ILEAL SODIUM- DEPENDENT BILE ACID TRANSPORTER) (ISBT) (SODIUM/TAURO CHOLATE COTRANSPORTIN G POLYPEPTIDE, ILEAL)	347	116/279 (41%)	173/279 (61%)	1e-52	
gi 8394281 ref NP 0 58918.1 NP 035518	solute carrier family 10, member 2 [Rattus norvegicus]	348	130/344 (37%)	195/344 (55%)	2e-52	
gi 6755530 ref NP 0 35518.1 NP 35518	solute carrier family 10, member 2 [Mus musculus]	348	125/313 (39%)	191/313 (60%)	4e-52	
gi 6755530 ref NP_0 35518.1 NP_35518	solute carrier family 10, member 2 [Mus musculus]	348	125/313 (39%)	191/313 (60%)	4e-52	

Applicants: U.S.S.N.:

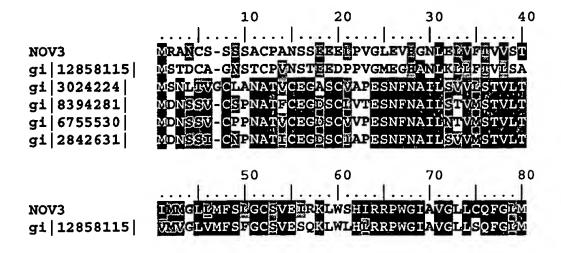
Guo 09/981,151

	,				
gi 2842631 sp Q6041	ILEAL	348	121/306	185/306	4e-52
4 NTCI CRIGR	SODIUM/BILE		(39%)	(59%)	
Q60414	ACID				ļ
	COTRANSPORTER				
	(ILEAL				Į.
	NA(+)/BILE				1
	ACID				ļ
ļ	COTRANSPORTER				
ł) (NA+				
·	DEPENDENT				
	ILEAL BILE				
	ACID				
	TRANSPORTER)				
	(ILEAL				
	SODIUM-				
	DEPENDENT				
	BILE ACID				
	TRANSPORTER)				
	(ISBT)				
	(SODIUM/TAURO		i		
	CHOLATE				
	COTRANSPORTIN				
	G				
	POLYPEPTIDE,				
	ILEAL)				

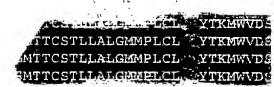
At page 53 please replace Table 3F with the following:

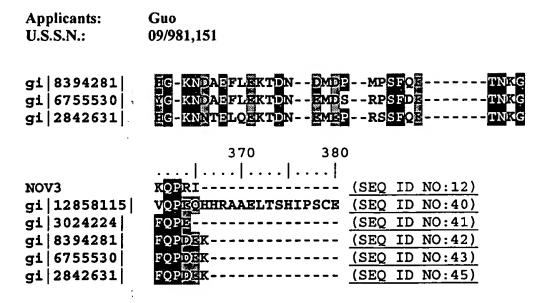
Table 3F. ClustalW Analysis of NOV3

- 1) NOV3 (SEQ ID NO:12)
- 3) gi|12858115|dbj|BAB31203.1| gi|12858115| BAB31203 (AK018423) putative [Mus musculus] (SEQ ID NO:40)
- 4) gi|3024224|sp|Q28727|NTCI RABIT g1|3024224| NP058918 ILEAL SODIUM/BILE ACID COTRANSPORTER (ILEAL NA(+)/BILE ACID COTRANSPORTER) (NA+ DEPENDENT ILEAL BILE ACID TRANSPORTER) (ILEAL SODIUM-DEPENDENT BILE ACID TRANSPORTER) (ISBT) (SODIUM/TAUROCHOLATE COTRANSPORTING POLYPEPTIDE, ILEAL) (SEQ ID NO:41)
- 5) <u>gi|8394281|ref|NP 058918.1| gi|8394281| NP035518</u> solute carrier family 10, member 2 [Rattus norvegicus] (SEQ ID NO:42)
- 6) gi|6755530|ref|NP-035518.1| gi|6755530| NP035518 solute carrier family 10, member 2 [Mus musculus] (SEQ ID NO:43)
- 7) <u>gi|6755530|ref|NP 035518.1|</u> <u>gi|6755530| NP035518</u> solute carrier family 10, member 2 [Mus musculus] (SEQ ID NO:44)
- 8) <u>gi|2842631|sp|Q60414|NTCI_CRIGR</u> gi|2842631| Q60414 ILEAL SODIUM/BILE ACID COTRANSPORTER (ILEAL NA(+)/BILE ACID COTRANSPORTER) (NA+ DEPENDENT ILEAL BILE ACID TRANSPORTER) (ILEAL SODIUM-DEPENDENT BILE ACID TRANSPORTER) (ISBT) (SODIUM/TAUROCHOLATE COTRANSPORTING POLYPEPTIDE, ILEAL) (SEQ ID NO:45)



Applicants: Guo **U.S.S.N.:** 09/981,151 ILLALVMFSMGCNVEI<mark>R</mark>KFLGHIRRPWGI<mark>F</mark>IGFLCOFGIM ILLALVMFSMGCNVEINKFLGHIRRPWGIFVGFLCOFGIM ILLALVMFSMGCNVEINKFLGHIRRPWGIFVGFLCOFGIM ILLALVMFSMGCNVEINKFLGHIRRPWGIVVGFLCOFGIM gi | 3024224 | gi | 8394281 | gi | 6755530 | gi | 2842631 | 100 PETAWILWISESIKPVQAWAVLIMGCCRG NOV3 gi | 12858115 | gi|3024224| PLTGFTLSVASGIPPVQAVVVLIMGCCPGGTGSNIL PLTGFTLSVASGIPPVQAVVVLIMGCCPGGTGSNIL PLTGFTLSVAFGIPPVQAVVVLIOGCCPGGTASNIL gi | 8394281 | gi|6755530| gi | 2842631 | 130 150 160 SGLMSTWISG-----ALGMMPLCLYSTYTWSWSLQQNTT
DGDMDLSTSMTTCSTLLALGMMPLCLYSTYTKMWVDSGTIV
DGDMDLSVSMTTCSTLLALGMMPLCLYSTYTKMWVDSGTIV
DGDMDLSVSMTTCSTLLALGMMPLCLSTYTKMWVDSGTIV
DGDMDLSVSMTTCSTLLALGMMPLCLSTYTKMWVDSGTIV
DGDMDLSVSMTTCSTLLALGMMPLCLSTYTKMWVDSGTIV NOV3 gi | 12858115 | gi|3024224| gi | 8394281 | gi|6755530| gi | 2842631 | 180 200 NOV3 gi | 12858115 | gi|3024224| IPYD----SIGISLVALVIPVSIGMÄVNHKWPOKAKITIK IPYD----SIGISLVALVIPVSIGMÄVNHKWPOKAKITIK IPYD----SIG<mark>I</mark>SLVALVIPVSIGMÄVNHKWPOKAKITIK gi | 8394281 | gi | 6755530 | gi 2842631 240 NOV3 gi | 12858115 | VGSIAGAVLIVLTÁVŘEGILYOSAWIIEPKĽWIIGTIEPÝ GSIAGAHLIVLIAVVČEILYOSAWIIEPKĽWIIGTIEP GSIIGVELIVLIAVEGGILYOSAWIIEPKĽWIIGTIEPE GSIAGATLÍVLIAVVGGILYOSAWIIEPKĽWIIGTIEPE gi 3024224 gi | 8394281 | gi | 6755530 | gi | 2842631 | 260 280 IGHVTGFLLALFTHOSWO--RTHPHFLGHAFKTPCDTHAAVGEVTGFLLAFFTHOSWORCRTHSHETGAONIOLCIAMO AGYSLGFFLARHAGOPWYRCRTVALETGYONTOLCSTIVO AGYSLGFFLARHAGOPWYRCRTVALETGYONTOLCSTIVO AGYSLGFFLARHAGOPWYRCRTVALETGYONTOLCSTIVO AGYGLGFFLARHAGOPWYRCRTVALETGYONTOLCSTIVO NOV3 gi | 12858115 | gi 3024224 gi | 8394281 | gi|6755530| gi | 2842631 | MSCPECSRLEGAFIPLEYGEFOREDGFETERE-LSFSAEYLVOELNEA-LAYGEFORENGLETVAAYQAYLSFSPEDLTYVFTFP-LIYSUFORVFAAGILGMYVTYLLSFSPEDLNLVFTFP-LIYBUFORVFAAGILGMYVTYLLSFSPEDLNLVFTFP-LIYBUFORVFAAGILGAYVAYLLSFSPEDLNLVFTFP-LIYBUFORAGILGAYVAYLLSFSPEDLNLVFTFP-LIYSUFORAGILGAYVAY NOV3 gi | 12858115 | gi|3024224| gi | 8394281 | gi|6755530| gi | 2842631 | 350|..<u>.</u>.|<u>.</u>...|.<u>.</u>..| ----RTEDTEC-----DGSP--LPEZFT----EVTIIP QKSKCRRÖHPDCPDVCYEKQPRETSAELDKGDEAAVTLGP EG-KNDAEFPDIKDT--KTEP--ESSFHQ------MNGG NOV3 gi | 12858115 | gi 3024224





Page 55, please replace Table 3G with the following

Table 3G Domain Analysis of NOV3

gnl|Pfam|pfam01758, SBF, Sodium Bile acid symporter family. This family consists of Na+/bile acid co-transporters. These transmembrane proteins function in the liver in the uptake of bile acids from portal blood plasma a process mediated by the co-transport of Na+. Also in the family is ARC3 from S. cerevisiae this is a putative transmembrane protein involved in resistance to arsenic compounds. (SEQ ID NO:82) CD-Length = 186 residues, 80.1% aligned Score = 75.9 bits (185), Expect = 3e-15

```
TIMMGLLMFSLGCSVEIRKLWSHIRRPWGIAVGLLCQFGLMPFTAYLLAIS-FSLKPVQA
Query:
                                  +||| + +||| |+ +|| ++||
             + + |+|||+|
                              ALGLFLMMFSMGLKVRFEDLKEALRRPKALILGLLLQWIIMPLLMFILAWLLLRLPPELA
Sbjct:
       1
Query:
       98
            IAVLIMGCCRG---APSLTFSPSGLMEIWIS-----GALGMMPLCIYLYTWSWSLQQN
              ++++|| +
                             +
                                 | +|+ +
                                                    + | | + |
            TGLILVGCAPGGAMSNVWTYLAKGDVELSVVMVALSTLLAPLVTPLLSFLLAGLL----
Sbjct:
       61
            LTIPYQNIGLSLGITLVCLTIPVAFGVYVNYRWP 181 (of SEQ ID NO:12)
Query:
                      + || + ||+ |+
       116 VHVDAVSPWSLIKSVLVYVIIPLIAGMLTRYFLP
Sbjct:
                                              149
                                                   (SEQ ID NO 82)
```

At page 61, please replace Table 4C with the following:

Table 4C. BLAST results for NOV4						
Gene Index/ Identifier	Protein/ Organism	Length (aa)	Identity (%)	Positives (%)	Expect	
gi 4505773 ref NP 0 02625.1 NP 002625	prohibitin [Homo sapiens]	272	236/270 (87%)	251/270 (92%)	e-123	
gi 92643 pir A3968 2 A39682	prohibitin - rat	272	235/270 (87%)	251/270 (92%)	e-123	

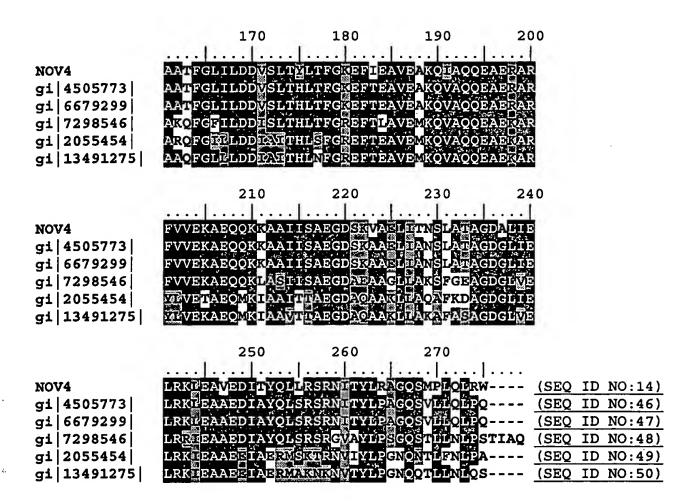
NOV4

gi | 4505773 | gi|6679299| gi | 7298546 | gi 2055454 gi | 13491275 |

gi 7298546 gb AAF53 765.1 (AE003661) AAF53765	l(2)37Cc gene product [Drosophila melanogaster]	276	178/270 (65%)	220/270 (80%)	8e-96
gi 2055454 gb AAB53 231.1 (U97204) AAB53231	prohibitin-like molecule TC-PRO-1 [Toxocara canis]	274	153/264 (57%)	209/264 (78%)	9e-80
gi 13491275 gb AAK2 7865.1 (AC087079) AAK27865	Hypothetical protein Y37E3.9 [Caenorhabditis elegans]	275	154/270 (57%)	210/270 (77%)	6e-79

At page 61 please replace Table 4D with the following:

Table 4D ClustalW Analysis of NOV4 1) NOV4 (SEQ ID NO:14) <u>gi | 4505773 | ref | NP 002625.1 | NP 002625 gi | 4505773 |</u> prohibitin [Homo sapiens] (SEQ 2) ID NO:46) gi|92643|pir||A39682 A39682 gi|6679299| prohibitin - rat (SEQ ID NO:47) 3) gi | 7298546 | gb | AAF53765.1 | AAF53765 (AE003661) gi | 7298546 | 1(2)37Cc gene product [Drosophila melanogaster] (SEQ ID NO:48) 5) gi|2055454|gb|AAB53231.1| AAB53231 (U97204) gi|2055454| prohibitin-like molecule TC-PRO-1 [Toxocara canis] (SEQ ID NO:49) 6) <u>gi|13491275|gb|AAK27865.1|</u> AAK27865 (AC087079) <u>gi|13491275|</u> Hypothetical protein Y37E3.9 [Caenorhabditis elegans] (SEQ ID NO:50) NOV4 gi | 4505773 | gi | 6679299 maa<mark>o fenr</mark>i go<mark>v</mark>gl<mark>gvavil</mark>ggvvnsályn gi | 7298546 | gi | 2055454 | MAASAOKULGRUGTVGVGLSEAGGUAQUALYNVI gi|13491275| 50 NOV4 gi | 4505773 | gi | 6679299 | Œ<mark>bD</mark>AAGEGIĤE<mark>E</mark> TSWAO gi | 7298546 | gi | 2055454 | IFDRF<mark>S</mark>GV<u>RE</u>VVGEGTHFLIPWVQKPIIFD<mark>I</mark>RS gi | 13491275 | 90 100 110 120 NOV4 gi | 4505773 | gi|6679299| gi | 7298546 | TITGSKOLONVŠITLRILHRPEP gi | 2055454 | TITGSKOLQNVNITLRILHRPSPDRLPNIY gi|13491275|



At page 62 please replace Table 4E with the following:

Table 4E. Domain Analysis of NOV4

gnl|Pfam|pfam01145, Band_7, SPFH domain / Band 7 family. This family
also includes proteins with high blast scores to known Band 7 protein:
HflC from E. coli HflK from E. coli, and Prohibitin family members
(SEQ ID NO:83)

CD-Length = 191 residues, 91.6% aligned Score = 157 bits (397), Expect = 7e-40

```
YSVDAGHRAVVFDRFRGVQDIVVGKGTHCLIPWLQKSIIFDCRSQPRNVPVITGSKDLQN
Query:
       28
                   1 1+1
                         | |+ ||| | +||++
                                                   | |+ +||
            KIVKEYERGVIFRLGRYVRQ-VVGPGLHFIIPFIDTVKKVDLRTVVYDVPSQEIITKDNV
Sbjct:
       17
            VNLTLRIIFRPVASQLPHIFTSSGEDHDERVPPSITNKILKSVVARFEAGELITQREQIS
Query:
       88
                +++ | | ++
                                        |++|+ +|
                                                            |++|+||+|+
            VVIVDAVVYYRVVDPLKAVYEVED---AERALPQLAQTTLRNVIGQFTLDEILTERERIN
Sbjct:
       76
                                                                         132
            {\tt RQVSDDLTEPAATFGLILDDVSLTYLTFGKEFIEAVEAKQIAQQEAERARFVVEKAEQQK}
Query:
       148
                         +|+ ++ | + + + | |+ |+ |+|| | |+ + + | |+
            SQLREILDEATDPWGIKVERVEIKDIRLPEEVQRAMAAQMEAEREA-RAKILEAEGEQEA
Sbjct:
Query: (aa 28-207 of SEQ ID NO:14)
```

Sbjct: (SEQ ID NO:83)

At page 63, please replace Table 4F with the following:

Table 4F. Domain Analysis of NOV4

gnl|Smart|smart00244, PHB, prohibitin homologues; prohibitin
homologues (SEQ ID NO:84)
CD-Length = 160 residues, 98.8% aligned
Score = 97.4 bits (241), Expect = 9e-22

```
YSVDAGHRAVVFDRFRGVQDIVVGKGTHCLIPWLQKSIIFDCRSQPRNVPVITG-SKDLQ
Query: 28
                            | |+| | | +||++
                      | +|
                                                   | |+| +||
Sbjct: 3
            FYVIGEGERGVVERLGRVLK-VLGPGLHFVIPFIDDVKRVDLRAQTDDVPPQEVITKDNV
            NVNLTLRIIFRPVASQLPHIFTSSGEDHDERVPPSITNKILKSVVARFEAGELIT-QREQ
Query:
                                     , | | |
                                                   |+||+ +
            TVSVDAVVYYR-VLDPLKAVYGV--LDADYRALRQLAQTTLRSVIGKRTLDELLTDEREK 118
Sbjct:
       62
           ISRQVSDDLTEPAATFGLILDDVSLTYLTFGKEFIEAVEAKQ
                                                       187 (of SEQ ID NO:14)
Query:
      146
            || + ++| | | +|+ ++|| + +
                                          +| ||+||+|
            ISENIREELNEAAEPWGIEVEDVEIKDIRLPEEIKEAMEAQQ
                                                       160 (SEQ ID NO:84)
Sbjct: 119
```

At page 67 please replace Table 5C with the following:

Table 5C. BLAST results for NOV5						
Gene Index/ Identifier	Protein/ Organism	Length (aa)	Identity (%)	Positives (%)	Expect	
gi 15294659 ref XP 054070.1 XP 05407	macrophage stimulating 1 (hepatocyte growth factor- like) [Homo sapiens]	711	561/720 (77%)	577/720 (79%)	0.0	
gi 10337615 ref NP 066278.1 NP 066278	macrophage stimulating 1 (hepatocyte growth factor- like) [Homo sapiens]	711	560/720 (77%)	576/720 (79%)	0.0	
gi 123114 sp P26927 HGFL_HUMAN_P26927	HEPATOCYTE GROWTH FACTOR-LIKE PROTEIN PRECURSOR (MACROPHAGE STIMULATORY PROTEIN) (MSP) (MACROPHAGE STIMULATING PROTEIN)	711	560/720 (77%)	576/720 (79%)	0.0	
gi 15299258 ref XP 017811.2 XP 017811	hypothetical protein XP_017811 [Homo sapiens]	529	440/532 (82%)	456/532 (85%)	0.0	
gi 90615 pir A4033 2 A40332	macrophage- stimulating protein 1 precursor - mouse	716	449/725 (61%)	507/725 (69%)	0.0	

Applicants:

Guo 09/981,151

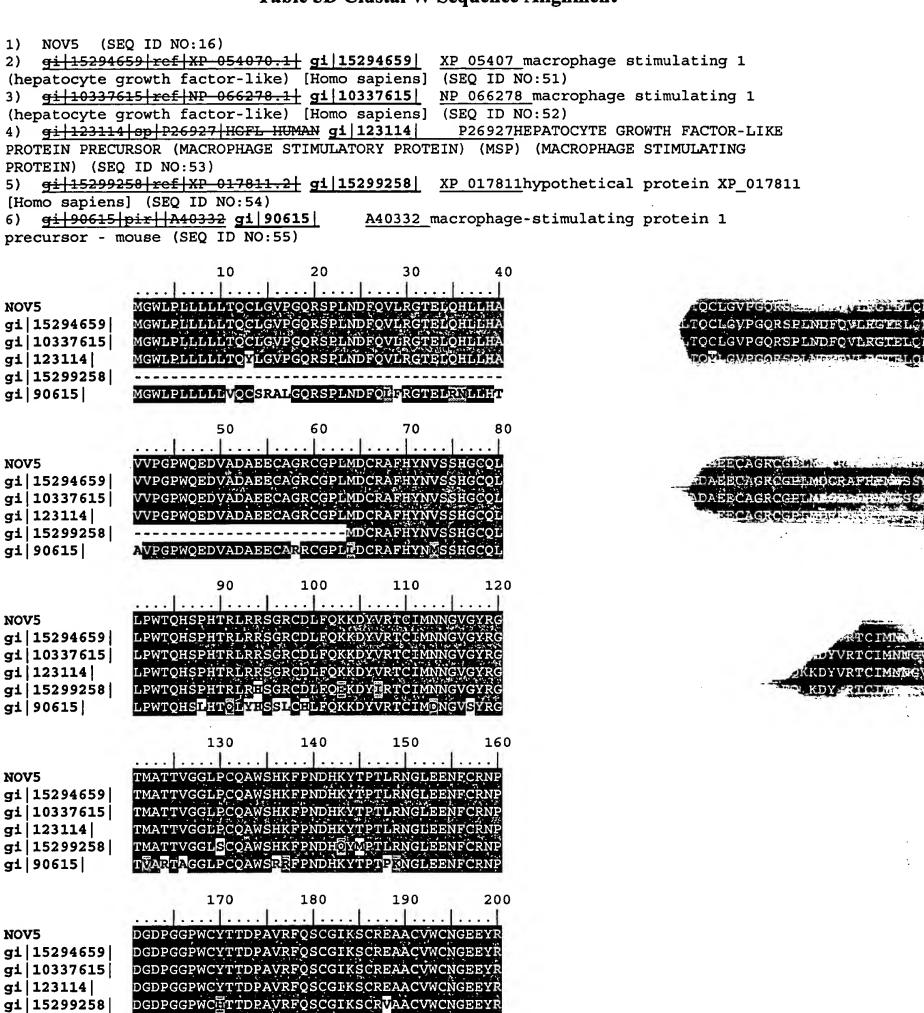
U.S.S.N.:

gi | 15299258 |

gi | 90615 |

At page 68 please replace Table 5D with the following:

Table 5D Clustal W Sequence Alignment



230

240

DGDPRGPWCYTTKRSVRFQSCGIKBCREAVCVLCNGEBYR

220

210

U.S.S.N.: 09/981,151 NOV5 GAVDRIESGRECORWDLOHPHOHPFEPGRFLDOGLDDNYC GAVDRIESGRECORWDLOHPHOHPFEPGRFLDOGLDDNYC GAVDRIESGRECORWDLOHPHOHPFEPGRFLDOGLDDNYC GAVDRIESGRECORWDLOHPHOHPFEPGRFLDOGLDDNYC GAVDRIESGRECORWDLOHPHOHPFEPGRFLDOGLDDNYC GEVDVITESGRECORWDLOHPHOHPFEPGRFLDOGLDDNYC gi | 15294659 | gi | 10337615 | gi | 123114 | gi | 15299258 | gi | 90615 | RNPDGSEREWCYTTDPOIEREFCDLPRCG RNPDGSEREWCYTTDPOIEREFCDLPRCG NOV5 gi | 15294659 | RNPDGSERPWCYTTDPOIEREFCDLPRCG----SE RNPDGSERPWCYTTDPOIEREFCDLPRCG---SE RNPDGSERPWCYTTDPOIEREFCDLPRCG---SE RNPDGSERPWCYTTDPNGEREFCDLPSCGPNLPPTVKGS gi|10337615| gi|123114| gi | 15299258 | gi | 90615 | NOV5 gi | 15294659 | gi | 10337615 | AOPROEATTYSEFRGKGEGYRGTANTITÄGVPCORV AOPROEATSVSCFRGKGEGYRGTANTITÄGVPCORV SORRNYGKANNEFRGKGE<mark>D</mark>YRGT<mark>T</mark>NTTSAGVPCORV gi | 123114 | g1 | 15299258 | gi|90615| PHOHRFTPEKYACKDLRENFCRNPDGSEAPWCFTLRPGMR
PHOHRFTPEKYACKDLRENFCRNPDGSEAPWCFTLRPGMR
PHOHRFTPEKYACKDLRENFCRNPDGSEAPWCFTLRPGMR
PHOHRFTPEKYACKDLRENFCRNPDGSEAPWCFTLRPGMR
PHOHRFTPEKYACKDLRENFCRNPDGSEAPWCFTLRPGMR
PHOHRFTPEKYACKDLRENFCRNPDGSEAPWCFTBRPGMR NOV5 gi | 15294659 | gi | 10337615 | gi | 123114 | gi | 15299258 | gi | 90615 | 380 400 AAFCYQIRRCTDDVRPQTATTAQGSSTAARSARPALVSSA
AAFCYQIRRCTDDVRPQDCYHGAGEQYRGTVSKTRKGVQC
AAFCYQIRRCTDDVRPQDCYHGAGEQYRGTVSKTRKGVQC
AAFCYQIRRCTDDVRPQDCYHGAGEQYRGTVSKTRKGVQC
VGFCYQIRRCTDDVRPQDCYHGAGEQYRGTVSKTRKGVQC
WAFCEQIPRCTGTVPECCYHGAGEQYRGTVSKTRKGVQC NOV5 gi | 15294659 | gi|10337615| gi | 123114 | gi | 15299258 | gi | 90615 | 420 | | | | | | . . <u>.</u> . NOV5 SAGPLRRRTSRSSRLPPNRMHNWRRTSAGTQMGIAMGPGA QRWSAETPHKPOFTFTSEPHAQLEENFCRNPDGDSHGP -QRWSAETPHKPOFTFTSEPHAQLEENFCRNPDGDSHGP -QRWSAETPHKPOFTFTSEPHAQLEENFCRNPDGDSHGP -QRGSAETPHKPOFTFTSEPHAQLEENFCTOMGTAMGPGA
QHWSGETPHKPOFTPTSAPOAGLEANFCRNPDGDSHGP -gi | 15294659 | gi | 10337615 | gi | 123114 | gi | 15299258 | gi | 90615 | 450 460 470 TRMTQGEHSTTVPCDAALMTSRHQS--WTEQTRCSLRSVA NOV5 WCYTHDPRIPPDYCALRRCADDQPPSILDPPDQVQFEKCG
WCYTHDPRIPPDYCALRRCADDQPPSILDPPDQVQFEKCG
WCYTHDPRIPPDYCALRRCADDQPPSILDPPDQVQFEKCG
TRITQGPHSTTVPCDAALMTSRHQS--WTPQTECSLRSVA
WCYTHDPDILFDYCALGRCDDDQPPSILDPPDQVVFEKCG gi | 15294659 | gi | 10337615 | gi | 123114 | gi | 15299258 | gi | 90615 |

Applicants:

Guo

....

RGTANTTTAGVECOR

ANTITAGVECORY

RGTANTTTAGTEC

GYRGTANTTTAGVI

U.S.S.N.: 09/981,151 NOV5 RGWIGWISGVPSCAW GAIRATH GQSACGIG---K<mark>RVDRLDORR</mark>SKLRVVGGHPGNSPWTVSLRNRQGQH gi 15294659 K<mark>RVDRLDORR</mark>SKLRVVGGHPGNSPWTVSLRNROGOHFCGC K<mark>RVDRLDORRS</mark>KLRVVGGHPGNSPWTVSLRNROGOHFCGC gi | 10337615 | gi | 123114 | GWIGWISVVPSCAW GAIRATH GOSACGIGOGOHFCGG gi | 15299258 | gi | 90615 | 530 540 550 -----MPLTGYEVWLGTLFQNPQHGE NOV5 SLVKEQWILTAROCFSSCHMPLTGYEVWLGTLFQNPOHGE SLVKEQWILTAROCFSSCHMPLTGYEVWLGTLFQNPOHGE SLVKEQWILTAROCFSSCHMPLTGYEVWLGTLFQNPOHGE SLVKEQWILTAROCFSSCHMPLTGYEVWLGTLFQNPOHGE SLVKEQWWLTAROCTWSCHEPLTGYEVWLGTENQNPOHGE gi | 15294659 | gi | 10337615 | gi | 123114 | gi | 15299258 | gi | 90615 | 580 590 PSLORVPVAKMVCGPSGSQLVLLKLERSVTLNORVALICL
PSLORVPVAKMVCGPSGSQLVLLKLERSVTLNORVALICL
PSLORVPVAKMVCGPSGSQLVLLKLERSVTLNORVALICL
PSLORVPVAKMVCGPSGSQLVLLKLERSVTLNORVALICL
PGLORVPVAKMVCGPSGSQLVLLKLERSVTLNORVALICL
ALORVPVAKAVCGPSGSQLVLLKLERFVTLNORVALICL NOV5 gi | 15294659 | gi | 10337615 | gi | 123114 | gi | 15299258 | gi | 90615 | PPEWYVVPPGTKCEIAGWGETKGTGNDTVLNVALLNVISN NOV5 PPEWYVVPPGTKCEIAGWGETKGTGNDTVLNVA<mark>L</mark>LNVISN gi | 15294659 | PPEWYVVPPGTKCEJAGWGETKGTGNDTVLNVA<mark>F</mark>LNVIŠN gi | 10337615 | gi | 123114 | gi | 15299258 | ppe<mark>o</mark>yvvppgtkceiagwges<mark>i</mark>gt<mark>s</mark>n\tvl@va<mark>s</mark>\nvisn gi | 90615 | 660 680 NOV5 gi | 15294659 | QECNIKHRGRVZESEVCT GLIAPVGACEGDYGGP QECNIKHRG<mark>RVZ</mark>ESEVCT GLIAPVGACEGDYGGP gi | 10337615 | gi | 123114 | gi | 15299258 | QECN<mark>o</mark>kärg::::©eseect@glivvpvgacegdyggplacät gi | 90615 | 700 710 . [. . . .]] . . | . . . HNCWVLEGIIIPNRVCARS<mark>C</mark>WPAVFTRVSVFVDWIHKVMR NOV5 HNCWVLEGIIIPNRVCARSRWPAVFTRVSVFVDWIHKVMR gi | 15294659 | HNCWVLEGITTPNRVCAŘSRWPAVFTRVSVFVDWTHKVMF gi | 10337615 | HNCWVLEGIIIPNRVCARSRWPAVFTRVSVFVDWIHKVM gi | 123114 | gi | 15299258 | HECWVL@GUIIPNRVCARPRWPAGFTRVSVFVDWINKVM® gi | 90615 | NOV5 (SEQ ID NO:16) gi | 15294659 | (SEQ ID NO:51) gi|10337615| (SEQ ID NO:52) (SEQ ID NO:53) gi|123114| (SEQ ID NO:54) gi | 15299258 | (SEQ ID NO:55) gi|90615|

Applicants:

Guo



At pages 70, please replace Table 5E with the following:

Table 5E. Domain Analysis of NOV5

gnl|Pfam|pfam00051, kringle, Kringle domain. Kringle domains have been found in plasminogen, hepatocyte growth factors, prothrombin, and apolipoprotein A. Structure is disulfide-rich, nearly all-beta. (SEQ ID NO:85) CD-Length = 79 residues, 100.0% aligned

CD-Length = 79 residues, 100.0% aligned Score = 117 bits (292), Expect = 3e-27

Query:	191			LQHPHQHPF-EPGKFLDQGLDDNYCRNPDGSERP	249
Sbjct:	. 1			SQTPHRHSKYTPERYPAKGLGENYCRNPDGDERP	60
Query:	250	WCYTTDPQIEREFCDLPRC 2	268	(of SEQ ID NO:16)	
Sbjct:	61	WCYTTDPRVRWEYCDIPRC	79 <u>(</u>	SEQ ID NO:85)	

At pages 71, please replace Table 5F, with the following:

Table 5F. Domain Analysis of NOV5

gnl|Pfam|pfam00051, kringle, Kringle domain. Kringle domains have been
found in plasminogen, hepatocyte growth factors, prothrombin, and
apolipoprotein A. Structure is disulfide-rich, nearly all-beta. (SEQ
ID NO:85)

CD-Length = 79 residues, 100.0% aligned Score = 112 bits (279), Expect = 9e-26

At pages 71, please replace Table 5G with the following:

Table 5G. Domain Analysis of NOV5

gnl|Pfam|pfam00051, kringle, Kringle domain. Kringle domains have been found in plasminogen, hepatocyte growth factors, prothrombin, and apolipoprotein A. Structure is disulfide-rich, nearly all-beta. (SEQ ID NO:85)

CD-Length = 79 residues, 100.0% aligned Score = 104 bits (259), Expect = 2e-23

At pages 71-72, please replace Table 5H with the following:

Table 5H. Domain Analysis of NOV5

gnl|Smart|smart00130, KR, Kringle domain; Named after a Danish pastry.
Found in several serine proteases and in ROR-like receptors. Can occur
in up to 38 copies (in apolipoprotein(a)). Plasminogen-like kringles
possess affinity for free lysine and lysine- containing peptides.
(SEQ ID NO:86)
CD-Length = 83 residues, 97.6% aligned

CD-Length = 83 residues, 97.6% aligned Score = 111 bits (278), Expect = 1e-25

At pages 72, please replace Table 5I with the following:

Table 5I. Domain Analysis of NOV5

gnl|Smart|smart00130, KR, Kringle domain; Named after a Danish pastry. Found in several serine proteases and in ROR-like receptors. Can occur in up to 38 copies (in apolipoprotein(a)). Plasminogen-like kringles possess affinity for free lysine and lysine- containing peptides. (SEQ ID NO:86)

CD-Length = 83 residues, 97.6% aligned Score = 106 bits (265), Expect = 4e-24

At pages 72, please replace Table 5J with the following:

Table 5J. Domain Analysis of NOV5

gnl|Smart|smart00130, KR, Kringle domain; Named after a Danish pastry.
Found in several serine proteases and in ROR-like receptors. Can occur
in up to 38 copies (in apolipoprotein(a)). Plasminogen-like kringles
possess affinity for free lysine and lysine- containing peptides.
(SEQ ID NO:86)
CD-Length = 83 residues, 97.6% aligned
Score = 104 bits (260), Expect = 1e-23

y: 283 CFRGKGEGYRGTANTTTAGVPCQRWDAQIPHQHRFTPEKYACKDLRENFCRNPDG-SEAP 34

Query: 342 WCFTLRPGMRAAFCYQIRRCTD 363 (of SEQ ID NO:16)

||+| | +| +| |+|

Sbjct: 63 WCYTTDPNVRWEYCD-IPQCES 83 (SEQ ID NO:86)

At pages 72-73, please replace Table 5K with the following:

Table 5K. Domain Analysis of NOV5

gnl|Smart|smart00020, Tryp_SPc, Trypsin-like serine protease; Many of these are synthesised as inactive precursor zymogens that are cleaved during limited proteolysis to generate their active forms. A few, however, are active as single chain molecules, and others are inactive due to substitutions of the catalytic triad residues. (SEQ ID NO:87) CD-Length = 230 residues, 79.1% aligned Score = 110 bits (274), Expect = 3e-25

```
504 PLTGYEVWLGTLFQNPQHGEPSLQRVPVAKMVCGPSGSQ-----LVLLKLERSVTLNQR 557
Query:
               | | | + +
                            | | |+|++ |+ +
                                              + | | | | |
                                                    |||+
          APSSIRVRLGSHDLSSGEET---QTVKVSKVIVHPNYNPSTYDNDIALLKLSEPVTLSDT
Sbjct:
      49
Query:
          VALICLPPEWYVVPPGTKCEIAGWGETKGTG--NDTVLNVALLNVISNQECNIKHRGR--
      558
          VRPICLPSSGYNVPAGTTCTVSGWGRTSESSGSLPDTLQEVNVPIVSNATCRRAYSGGPA
Sbjct:
      106
                                                            165
            Query:
      614
                         ||| ||+
                                               | |+|||
          ITDNMLCAGGLEGGKDACQGDSGGPLVC-NDPRWVLVGIVSWGSYGCARPNKPGVYTRVS
Sbjct:
      166
Query:
      654
          VFVDWI
                659
                    (of SEQ ID NO:16)
           ++|||
Sbjct:
      225 SYLDWI
                230
                    (SEQ ID NO:87)
```

At pages 73, please replace 5L with the following:

Table 5L. Domain Analysis of NOV5

gnl|Pfam|pfam00089, trypsin, Trypsin. Proteins recognized include all
proteins in families S1, S2A, S2B, S2C, and S5 in the classification
of peptidases. Also included are proteins that are clearly members,
but that lack peptidase activity, such as haptoglobin and protein Z
(PRTZ*). (SEQ ID NO:88)
CD-Length = 217 residues, 77.9% aligned
Score = 92.0 bits (227), Expect = 9e-20

```
YEVWLGTLFQNPQHGEPSLQRVPVAKMV----CGPSGSQLVLLKLERSVTLNQRVALICL
Query:
                     | + |+ | |++
                                           | + + ||||+ |||
            VRVVLGE--HNLGTTEGTEQKFDVKKIIVHPNYNPDTNDIALLKLKSPVTLGDTVRPICL
Sbjct:
       49
                                                                      106
           PPEWYVVPPGTKCEIAGWGETKGTGNDTVLNVALLNVISNQECNIKHRG------
Query:
                 +| || | ++||| || |
                                          ++ ++| + | + |
            PSASSDLPVGTTCSVSGWGRTKNLGTSDTLQEVVVPIVSRETCRSAYGGTVTDTMICAGA 166
       107
Sbjct:
            -----RGDYGGPLACFTHNCWVLEGIIIPNRVCARSCWPAVFTRVSVFVDWI 659
Query:
       613
                  +|| ||| |
                                  | ||+
                                          +| |+||| ++||
Sbjct: 167 LGGKDACQGDSGGPLVCSDG---ELVGIVSWGYGCAVGNYPGVYTRVSRYLDWI
```

Query: (of SEQ ID NO:16)
Sbjct: (SEQ ID NO:88)

At pages 73, please replace Table 5M with the following:

Table 5M. Domain Analysis of NOV5

gnl|Smart|smart00473, PAN_AP, divergent subfamily of APPLE domains; Apple-like domains present in Plasminogen, C. elegans hypothetical ORFs and the extracellular portion of plant receptor-like protein kinases. Predicted to possess protein- and/or carbohydrate-binding functions. (SEQ ID NO:89) CD-Length = 79 residues, 94.9% aligned Score = 52.0 bits (123), Expect = 1e-07

At pages 74, please replace Table 5N with the following:

Table 5N. Domain Analysis of NOV5

gnl|Pfam|pfam00024, PAN, PAN domain. The PAN domain contains a conserved core of three disulphide bridges. In some members of the family there is an additional fourth disulphide bridge the links the N and C termini of the domain. The domain is found in diverse proteins, in some they mediate protein-protein interactions, in others they mediate protein-carbohydrate interactions. (SEQ ID NO:159) CD-Length = 78 residues, 76.9% aligned Score = 50.1 bits (118), Expect = 4e-07

At page 78 please replace Table 6C with the following:

Applicants: U.S.S.N.:

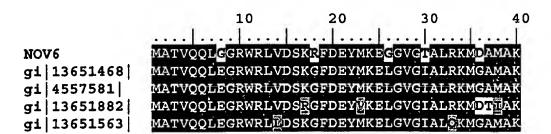
Guo 09/981,151

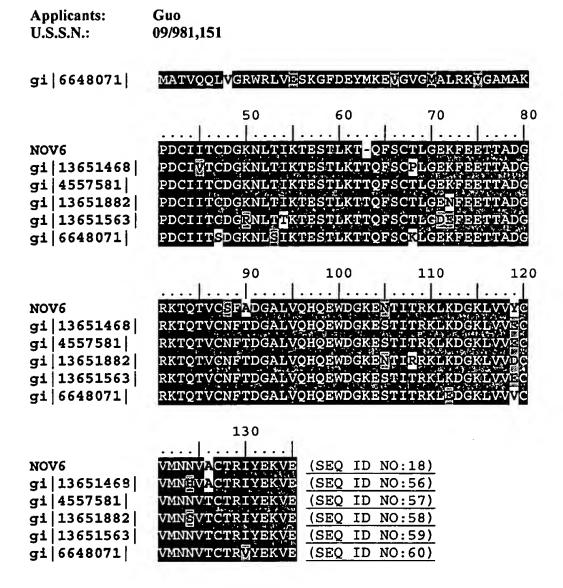
	Table 6C. BLA	ST resul	ts for NOV	6	
Gene Index/ Identifier	Protein/ Organism	Length (aa)	Identity (%)	Positives (%)	Expect
gi 13651468 ref XP 016351.1 XP 016351	similar to GASTRIN/CHOLECYST OKININ TYPE B RECEPTOR (CCK-B RECEPTOR) (CCK-BR) (H. sapiens) [Homo sapiens]	135	122/135 (90%)	126/135 (92%)	1e-58
gi 4557581 ref NP 0 01435.1 NP 001435	fatty acid binding protein 5 (psoriasis- associated); E- FABP [Homo sapiens]	135	124/135 (91%)	126/135 (92%)	1e-58
gi 13651882 ref XP 011655.5 XP 011655	fatty acid binding protein 5 (psoriasis- associated) [Homo sapiens]	135	119/135 (88%)	124/135 (91%)	6e-57
gi 13651563 ref XP- 015760.1 XP 015760	similar to GASTRIN/CHOLECYST OKININ TYPE B RECEPTOR (CCK-B RECEPTOR) (CCK-BR) (H. sapiens) [Homo sapiens]	135	118/135 (87%)	125/135 (92%)	5e-56
gi 6648071 sp P5505 2 FABE BOVIN P55052	FATTY ACID- BINDING PROTEIN, EPIDERMAL (E- FABP)	135	117/135 (86%)	124/135 (91%)	1e-55

At page 79 please replace Table 6D with the following:

Table 6D Information for the ClustalW proteins

- 1) NOV6 (SEQ ID NO:18)
- 2) gi|13651468|ref|XP 016351.1| gi|13651468| XP 016351similar to GASTRIN/CHOLECYSTOKININ TYPE B RECEPTOR (CCK-B RECEPTOR) (CCK-BR) (H. sapiens) [Homo sapiens] (SEQ ID NO:56)
- 3) gi|4557581|ref|NP 001435.1| gi|4557581| NP 001435 fatty acid binding protein 5 (psoriasis-associated); E-FABP [Homo sapiens] (SEQ ID NO:57)
- 4) <u>gi|13651882|ref|XP 011655.5|</u> <u>gi|13651882|</u> <u>XP 011655</u>fatty acid binding protein 5 (psoriasis-associated) [Homo sapiens] (SEQ ID NO:58)
- 5) <u>gi|13651563|ref|XP 015760.1|</u> <u>gi|13651563|</u> <u>XP 015760</u>similar to
- GASTRIN/CHOLECYSTOKININ TYPE B RECEPTOR (CCK-B RECEPTOR) (CCK-BR) (H. sapiens) [Homo sapiens] (SEQ ID NO:59)
- 6) gi | 6648071 | sp | P55052 | FABE BOVIN gi | 13651563 | P55052 | FATTY ACID-BINDING PROTEIN, EPIDERMAL (E-FABP) (SEQ ID NO:60)





Page 80, please replace Table 6E with the following:

Sbjct: (SEQ ID NO:90)

Table 6E. Domain Analysis of NOV6

gnl|Pfam|pfam00061, lipocalin, Lipocalin / cytosolic fatty-acid
binding protein family. Lipocalins are transporters for small
hydrophobic molecules, such as lipids, steroid hormones, bilins, and
retinoids. Alignment subsumes both the lipocalin and fatty acid
binding protein signatures from PROSITE. This is supported on
structural and functional grounds. Structure is an eight-stranded beta
barrel. (SEQ ID NO:90)
CD-Length = 145 residues, 76.6% aligned
Score = 53.1 bits (126), Expect = 1e-08

Applicants: Guo U.S.S.N.: 09/981,151

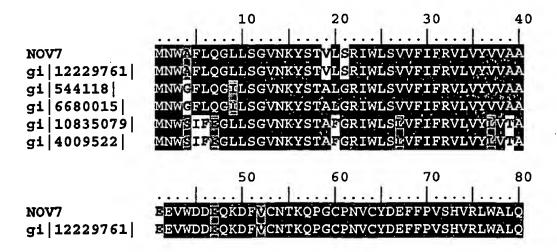
At page 84 please replace Table 7C with the following:

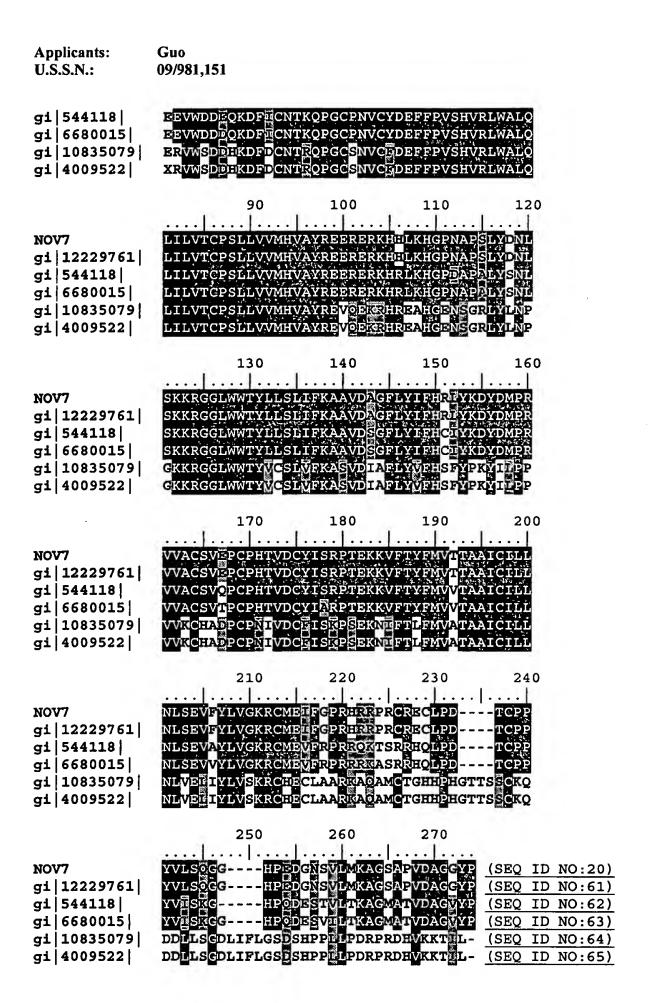
Table 7C. BLAST results for NOV7						
Gene Index/ Identifier	Protein/ Organism	Length (aa)	Identity (%)	Positives (%)	Expect	
gi 12229761 sp Q9NTQ 9 CXB4_HUMAN Q9NTQ9	GAP JUNCTION BETA-4 PROTEIN (CONNEXIN 30.3) (CX30.3)	266	266/266 (100%)	266/266 (100%)	e-155	
gi 544118 sp P36380 CXB5 RAT P36380	GAP JUNCTION BETA-5 PROTEIN (CONNEXIN 30.3) (CX30.3)	265	230/266 (86%)	245/266 (91%)	e-132	
gi 6680015 ref NP-03 2153.1 NP 032153	gap junction membrane channel protein beta 4; connexin 30.3 [Mus musculus]	266	230/266 (86%)	244/266 (91%)	e-131	
gi 10835079 ref NP-0 05259.1 NP 05259	gap junction protein, beta 5 (connexin 31.1) [Homo sapiens]	273	153/226 (67%)	177/226 (77%)	3e-88	
gi 4009522 gb AAC954 72.1 (AF099731)AAC95472	connexin 31.1 [Homo sapiens]	273	152/226 (67%)	176/226 (77%)	1e-87	

At page 85 please replace Table 7D with the following:

Table 7D. Information for the ClustalW proteins

- 1) NOV7 (SEQ ID NO:20)
- 2) gi | 12229761 | sp | Q9NTQ9 | CXB4 HUMAN Q9NTQ9 GAP JUNCTION BETA-4 PROTEIN (CONNEXIN 30.3) (CX30.3) (SEQ ID NO:61)
- 3) $\frac{gi|544118|sp|P36380|CXB5-RAT}{CX30.3}$ GAP JUNCTION BETA-5 PROTEIN (CONNEXIN 30.3) (CX30.3) (SEQ ID NO:62)
- 4) gi|6680015|ref|NP-032153.1| NP 032153gap junction membrane channel protein beta 4; connexin 30.3 [Mus musculus] (SEQ ID NO:63)
- 5) gi|10835079|ref|NP 005259.1| NP 05259 gap junction protein, beta 5 (connexin 31.1) [Homo sapiens] (SEQ ID NO:64)
- 6) <u>gi|4009522|gb|AAC95472.1|</u> AAC95472 (AF099731) connexin 31.1 [Homo sapiens] (SEQ ID NO:65)





At page 86 please replace Table 7E with the following:

Applicants: Guo U.S.S.N.: 09/981,151

Table 7E. Domain Analysis of NOV7

gnl|Pfam|pfam00029, connexin, Connexin. (SEQ ID NO:91)
CD-Length = 218 residues, 100.0% aligned
Score = 318 bits (814), Expect = 3e-88

Query:	1	MNWAFLQGLLSGVNKYSTVLSRIWLSVVFIFRVLVYVVAA				60
Sbjct:	1	MDWSFLGRLLEGVNKHSTAIGKIWLSVLFIFRILVLGVAA				60
Query:	61	PNVCYDEFFPVSHVRLWALQLILVTCPSLLVVMHVAYREE			SLYDNL	120
Sbjct:	61	ENVCYDQFFPISHVRLWVLQLIFVSTPSLLYLGHVAYRVR		•		120
Query:	121	SKKRGGLWWTYLLSLIFKAAVDAGFLYIF				169
Sbjct:	121	AKKRCGSEDGKVRIRGGLWWTYVFSIIFKSIFEVGFLYGQ				179
Query:	170	PHTVDCYISRPTEKKVFTYFMVTTAAICILLNLSEVFYL	208	(of SEQ	ID NO:2	(0)
Sbjct:	180	PHTVDCFVSRPTEKTIFIVFMLVVSAICLLLNLAELFYL	218	(SEQ ID	NO:91)	

At page 87, please replace Table 7F with the following:

Table 7F. Domain Analysis of NOV7

gnl|Smart|smart00037, CNX, Connexin homologues; Connexin channels
participate in the regulation of signaling between developing and
differentiated cell types. (SEQ ID NO:160)
CD-Length = 34 residues, 100.0% aligned
Score = 79.0 bits (193), Expect = 3e-16

At page 100, please replace Table 9C with the following:

Table 9C. BLAST results for NOV9					
Gene Index/ Identifier	Protein/ Organism	Length (aa)	Identity (%)	Positives (%)	Expect
gi 13591536 emb CAC 36351.1 CAC36351 (AL109613)	dJ1033H22.1 (KIAA0554 protein) [Homo sapiens]	434	373/430 (86%)	375/430 (86%)	0.0

Applicants: U.S.S.N.:

Guo 09/981,151

gi 8923249 ref NP 0 60207.1 NP 060207	hypothetical protein FLJ20275 [Homo sapiens]	330	328/330 (99%)	329/330 (99%)	e-175
gi 10435680 dbj BAB 14638.1 BAB14638 (AK023681)	unnamed protein product [Homo sapiens]	592	319/595 (53%)	432/595 (71%)	e-160
gi 13936547 gb AAK4 9824.1 AF265550 1 AAK49824 (AF265550)	formin-binding protein 17 [Homo sapiens]	679	307/624 (49%)	422/624 (67%)	e-148
gi 3043632 dbj BAA2 5480.1 BAA25480 (AB011126)	KIAA0554 protein [Homo sapiens]	674	307/624 (49%)	422/624 (67%)	e-148

At page 100 please replace Table 9D with the following:

Table 9D. ClustalW Analysis of NOV9

3) gi 135915 (KIAA0554 pro 4) gi 892324 [Homo sapiens 5) gi 104356 product [Homo 6) gi 139365 binding prote 7) gi 304363	9 (SEQ ID NO:24) 36 emb CAC36351.1 CAC36351 gi 13591536 (AL109613) dJ1033H22.1 tein) [Homo sapiens] (SEQ ID NO:66) 9 ref NP 060207.1 NP 060207 gi 8923249 hypothetical protein FLJ20275] (SEQ ID NO:67) 80 dbj BAB14638.1 BAB14638 gi 10435680 (AK023681) unnamed protein sapiens] (SEQ ID NO:68) 47 gb AAK49824.1 AF265550 AAK49824 gi 13936547 (AF265550) forminin 17 [Homo sapiens] (SEQ ID NO:69) 2 dbj BAA25480.1 BAA25480 gi 3043632 (AB011126) KIAA0554 protein] (SEQ ID NO:70)
NOV9 gi 13591536 gi 8923249 gi 10435680 gi 13936547 gi 3043632	10 20 30 40 LWNGGEEEPPRRPRARSCEPEEAARTPGFPPSRGSRGAKGEEEPPRRPRARSCEPEEAARTPGFPPSRGSRGAKG
NOV9 gi 13591536 gi 8923249 gi 10435680 gi 13936547 gi 3043632	50 60 70 80
NOV9 gi 13591536 gi 8923249 gi 10435680 gi 13936547 gi 3043632	90 100 110 120

Applicants: Guo **U.S.S.N.:** 09/981,151 130 140 160 NOV9 gi|13591536| gi | 8923249 | SKEEB<mark>EY W</mark>TSCKAF<mark>ISN</mark>LNE NDYAGOHEVIS SKEEB<mark>EY W</mark>TSCKAFISNLNE NDYAGOHEVIS SKEEB<mark>EY W</mark>TSCKAFISNLNE NDYAGOHEVIS gi | 10435680 | gi | 13936547 | gi|3043632| 200 NOV9 YGELMRYAHDLKTERKMHLQEGRKAQQYAADMCWKQYG\SK gi | 13591536 | gi | 8923249 | IVILARYVO ELKOERKS VITTOGRKAQQEGGETICWKO
IVILARYVO ELKOERKS VITTOGRKAQQEGGETICWKQ
IVILARYVO ELKOERKS VITTOGRKAQQEGGETICWKQ gi | 10435680 | gi | 13936547 | gi 3043632 220 240 230 NOV9 gi | 13591536 | gi | 8923249 | PRFERECKEADRAQOV JERODADINVTKADVEKAROOAQU RRFERECKEADRAQOV JERODADINVTKADVEKAROOAQU RRFERECKEADRAQOV JEROVDADINVTKADVEKAROOAQU gi|10435680| gi | 13936547 | gi 3043632 NOV9 gi | 13591536 | gi 8923249 RHOMAGDSKA YSSILOKENBEQHSAAGTHIPNIGOGOE RHOMAGDSKA YSSILOKENBEQHSAAGTHIPNIGOGOE RHOMAGDSKA YSSILOKENBEQHSAAGTHIPNIGOGOE gi | 10435680 | gi | 13936547 | gi 3043632 290 320 NOV9 gi | 13591536 | gi | 8923249 | gi | 10435680 | gi | 13936547 | gi | 3043632 | 350 340 360 NOV9 gi|13591536| gi | 8923249 | IDO MOSO V O KKŚĆĘĘPED IEFEDY OPK IDO MOSO V O KKŚĆĘĘPEDIEFEDY OPK IDO MOSO V O KKŚĆĘĘPEDIEFEDY OPK gi | 10435680 | gi | 13936547 | gi|3043632| 380 400 NOV9 gi | 13591536 | gi | 8923249 | KGKLW DLKFG-GKSKGKLWPFIKKN DLKFG-GKSKGKLWPFIKKNKLMSLLTS DLKFG-GKSKGKLWPFIKKNKLMSLLTS gi | 10435680 | gi | 13936547 |

GKPDLKFG.

410

420

g1|3043632|

430

440

-SINISTING WAR OF THE INTERIOR

U.S.S.N.: 09/981,151 | | | | | | | NOV9 --Qsppltptslftsstpngs@fltfsi<mark>epver</mark>cm<mark>ne</mark>ik<mark>t</mark> gi | 13591536 | gi|8923249| PHQPPPPPPASASPSAVPNGPOSPKQQKEPISERFNEFMT PHQPPPPPPASASPSAVPNGPOSPKQQKEPISERFNEFMT gi | 10435680 | gi | 13936547 | gi|3043632| 450 460 | | | | . . . NOV9 gi | 13591536 | gi | 8923249 | SKPKIHCFRSLKRGLSLKLGATPEDFS gi|10435680| SKPRIHOFRSLKRG<mark>LSLKLGATP</mark>EDFSNLPPEORRE SKPRIHOFRSLKRG<mark>LSLKLGATP</mark>EDFSNLPPEORRE gi|13936547| gi 3043632 NOV9 gi|13591536| OP gi|8923249| gi | 10435680 | DH WDELNGETOKEMDORDAETKMKDVYLKNEOMGDPASLDH WDELNGETOKEMDORDAETKMKDVYLKNEOMGDPASLDH gi|13936547|-9 DELN gi|3043632| 560 KLAETMINIDELEVEGIHKNEAWLSEVEG NOV9 KLAETMINNIDRLE gi | 13591536 | gi |8923249| gi | 10435690 | gi|13936547| KLAEVSONIEKLRYETOK FEAWLZEVEGRLPARSEOA gi|3043632| 570 580 590 600 NOV9 gi|13591536| gi|8923249| gi | 10435680 | gi | 13936547 | gi|3043632| 610 630 HGHHNEFDDEFEDDDPLPAIGHCKAMYPEDGHNEGTMANK
HGHHNEFDDEFEDDDPLPAIGHCKAMYPEDGHNEGTMANK
HGHHNEFDDEFEDDDPLPAIGHCKAMYPEDGHNEGTMANK
KVLATDFDDEFDDEEPLPAIGTCKAMYTFEGONEGTESVV
KVLATDFDDEFDDEEPLPAIGTCKAMYTFEGONEGTESVV
KVLATDFDDEFDDEEPLPAIGTCKAMYTFEGONEGTESVV NOV9 gi | 13591536 | gi|8923249| gi | 10435680 | gi | 13936547 | gi|3043632| 680 650 660 670 NOV9 EGEVLYMIEEDKGDGWTRARRONGEEGYVPTSYMEGEVLYMIEEDKGDGWTRARRONGEEGYVPTSYMEGETLYMIEEDKGDGWTRARRONGEEGYVPTSYMEGETLYMIEEDKGDGWTRARRONGEEGYVPTSYMEGETLYMIEEDKGDGWTRARRONGDEEGYVPTSYMEGETLYMIEEDKGDGWTRARRONGEEGYVPTSYMEGETLYMIEEDKGDGWTRARRONGEEGYVPTSYMEGEGTLYMIEEDKGDGWTRARRONGEEGYVPTSYMEGEGTLYMIEEDKGDGWTRARRONGEEGYVPTSYMEGEGTLYMIEEDKGDGWTRARRONGEEGYVPTSYMEGEGTLYMIEEDKGDGWTRARRONGEEGYVPTSYMEGEGTLYMIEEDKGDGWTRARRONGEEGYVPTSYMEGEGTVPTSYMEGETTSYMEGTTS gi|13591536| gi | 8923249 | gi|10435680| gi|13936547| gi|3043632|

. . . . |

Applicants:

Guo

U.S.S.N.: 09/981,151

NOV9
gi | 13591536 | NSKGAVTYI (SEQ ID NO:66)
gi | 8923249 | NSKGAVTYI (SEQ ID NO:67)
gi | 10435680 | NAKGAKTYI (SEQ ID NO:68)
gi | 13936547 | NAKOS---- (SEQ ID NO:69)
gi | 3043632 | NAKOS---- (SEQ ID NO:70)

Guo

Applicants:

At page 103, please replace Table 9E with the following:

Table 9E. Domain Analysis of NOV9

gnl|Smart|smart00326;—SH3, Src homology 3 domains; Src homology 3 (SH3) domains bind to target proteins through sequences containing proline and hydrophobic amino acids. Pro-containing polypeptides may bind to SH3 domains in 2 different binding orientations. (SEQ ID NO:92)

CD-Length = 59 residues, 88.1% aligned Score = 64.7 bits (156), Expect = 1e-11

Query: 484 HCKAIYPFDGHNEGTLAMKEGEVLYIIEEDKGDGWTRARRQNGEEGYVPTSYI 536

Sbjct: 4 QVRALYDYTAQDPDELSFKKGDIITVLEKS-DDGWWKGRLGTGKEGLFPSNYV 55

Query: (of SEQ ID NO:24)

Sbjct: (SEQ ID NO:92)

At page 104, please replace Table 9F with the following:

Table 9F. Domain Analysis of NOV9

gnl|Pfam|pfam00018, SH3, SH3 domain. SH3 (Src homology 3) domains are often indicative of a protein involved in signal transduction related to cytoskeletal organization. First described in the Src cytoplasmic tyrosine kinase. The structure is a partly opened beta barrel. (SEQ ID NO:93)

CD-Length = 57 residues, 91.2% aligned Score = 63.5 bits (153), Expect = 3e-11

Query: 486 KAIYPFDGHNEGTLAMKEGEVLYIIEEDKGDGWTRARRQNGEEGYVPTSYID 537

A+Y+ L+ K+G+++ ++E+ GW+ R+ +EG+P++Y++

Sbjct: 4 VALYDYQARESDELSFKKGDIIIVLEKSDDGGWWKGRLKGTKEGLIPSNYVE 55

Query: of SEQ ID NO:24)

Sbjct: (SEQ ID NO:93)

Applicants: Guo U.S.S.N.: 09/981,151

At page 104, please replace Table 9 with the following:

Table 9G. Domain Analysis of NOV9

gnl|Smart|smart00055, FCH, Fes/CIP4 homology domain; Alignment
extended from original report. Highly alpha-helical. Also known as the
RAEYL motif or the S. pombe Cdc15 N-terminal domain. (SEQ ID NO:94)
CD-Length = 91 residues, 97.8% aligned
Score = 58.2 bits (139), Expect = 1e-09

At page 104, please replace Table 9H with the following:

Table 9H. Domain Analysis of NOV9

gnl|Pfam|pfam00611, FCH, Fes/CIP4 homology domain. Alignment extended
from. Highly alpha-helical. (SEQ ID NO:95)
CD-Length = 94 residues, 97.9% aligned
Score = 40.0 bits (92), Expect = 3e-04

At page 113 please replace Table 10F with the following:

Applicants: U.S.S.N.:

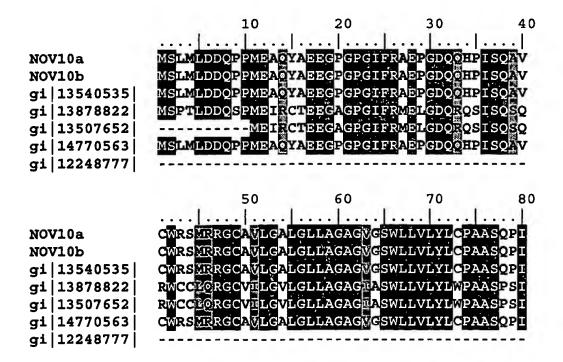
Guo 09/981,151

Table 10F. BLAST results for NOV10a						
Gene Index/ Identifier	Protein/ Organism	Length (aa)	Identity (%)	Positives (%)	Expect	
gi 13540535 ref NP 110397.1 NP 110397	transmembrane protease, serine 5; spinesin [Homo sapiens]	457	452/463 (97%)	453/463 (97%)	0.0	
gi 13878822 sp Q9ER 04 TM65 MOUSE Q9ER04	TRANSMEMBRANE PROTEASE, SERINE 5 (SPINESIN)	455	360/461 (78%)	394/461 (85%)	0.0	
gi 13507652 ref NP 109634.1 NP 109634	transmembrane protease, serine 5 (spinesin) [Mus musculus]	445	354/451 (78%)	388/451 (85%),	0.0	
gi 14770563 ref XP 041427.1 -XP 041427	transmembrane protease, serine 5 [Homo sapiens]	398	354/362 (97%)	355/362 (97%)	0.0	
gi 12248777 dbj BAB 20276.1 BAB20276 (AB016229)	type 2 spinesin [Mus musculus]	311	260/317 (82%)	281/317 (88%)	e-146	

At page 114 please replace Table 10G with the following:

Table 10G. ClustalW Analysis of NOV10

- 1) Novel NOV10a (SEQ ID NO:27)
- 2) Novel NOV10b (SEQ ID NO:29)
- 4) <u>gi|13540535|ref|NP-110397.1|</u> NP 110397 **gi|13540535|** transmembrane protease, serine 5; spinesin [Homo sapiens] (SEQ ID NO:71)
- 5) gi|13878822|sp|Q9ER04|TMS5-MOUSE Q9ER04 gi|13878822| TRANSMEMBRANE PROTEASE, SERINE 5 (SPINESIN) (SEQ ID NO:72)
- 6) <u>gi|13507652|ref|NP-109634.1|</u> NP 109634 **gi|13507652|** transmembrane protease, serine 5 (spinesin) [Mus musculus] (SEQ ID NO:73)
- 7) <u>gi|14770563|ref|XP 041427.1|</u> XP 041427 **gi|14770563|** transmembrane protease, serine 5 [Homo sapiens] (SEQ ID NO:74)
- 8) <u>gi|12248777|dbj|BAB20276.1|</u> BAB20276 <u>gi|12248777|</u> (AB016229) type 2 spinesin [Mus musculus] (SEQ ID NO:75)



120 NOV10a NOV10b gi | 13540535 | gi|13878822| gi | 13507652 | gi | 14770563 | gi | 12248777 | 160 NOV10a NOV10b gi|13540535| gi | 13878822 | gi | 13507652 | gi | 14770563 | gi | 12248777 | NOV10a NOV10b gi | 13540535 | DIKLNRSOEFAOLSA gi|13878822| gi | 13507652 | GVNLIDIKLN<mark>SSQEFAQLSPRLGGFTEEAW</mark> AVNLSDIKLNRSQEFAQLSARPGGLVEEAW gi | 14770563 | gi | 12248777 | 240 NOV10a NOV10b gi | 13540535 | gi | 13878822 | gi | 13507652 | gi | 14770563 | CSECGARPLASRIV SG VSL CSECGARPLASRIVGGO gi | 12248777 | NOV10a NOV10b gi|13540535| gi | 13878822 | gi | 13507652 | gi | 14770563 | gi|12248777| 290 320 NOV10a NOV10b HAGLVSHSAVRPHOGA EVERTIPHPLYSAONHDYDVALL gi | 13540535 | HAGLVSHGAVROHOGATVVEKI I PHPLYSAQNHDYDVÄLLOHAGLVSHGAVROHOGATVVEKI I PHPLYSAQNHDYDVÄLLOHAGLVSHGAVROHOGATVVEKI I PHPLYSAQNHDYDVALLOHAGLVSHGAVROHOGATVVEKI I PHPLYSAQNHDYDVALLOHAGLVSHGAVROHOGATVVEKI I PHPLYSAQNHDYDVALLO gi | 13878822 | gi | 13507652 | gi | 14770563 | gi | 12248777 | 360 330 NOV10a

Applicants:

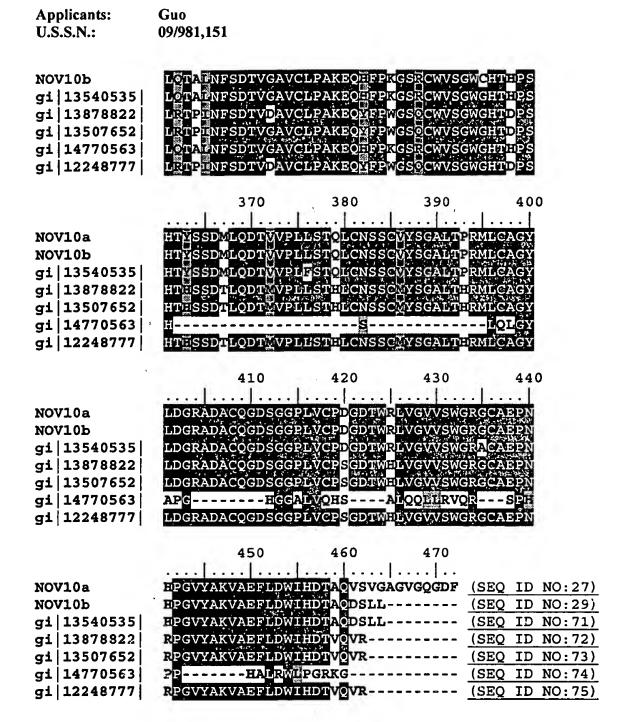
U.S.S.N.:

Guo

09/981,151











At page 116, please replace Table 10H with the following:

Table 10H Domain Analysis of NOV10

gnl|Smart|smart00020, Tryp_SPc, Trypsin-like serine protease; Many of these are synthesised as inactive precursor zymogens that are cleaved during limited proteolysis to generate their active forms. A few, however, are active as single chain molecules, and others are inactive due to substitutions of the catalytic triad residues. (SEQ ID NO:96) CD-Length = 230 residues, 100.0% aligned Score = 266 bits (681), Expect = 2e-72

```
Query:
       224
            RIVGGQSVAPGRWPWQASVAL-GFRHTCGGSVLAPRWVVTAAHCMHSFRLARLSSWRVHA
                      | +||| |+ || || |||+++||||+||||++
            RIVGGSEANIGSFPWQVSLQYRGGRHFCGGSLISPRWVLTAAHCVYGSAP---SSIRVRL
Sbjct:
            GLVSHSAVRPHQGALVERIIPHPLYSAQNHDYDVALLRLQTALNFSDTVGAVCLPAKEQH
Query:
       283
                           | ++| || |+
                                        +| |+|||+|
                                                     + |||| +|||+
            GSHDLSSGEETQTVKVSKVIVHPNYNPSTYDNDIALLKLSEPVTLSDTVRPICLPSSGYN
Sbjct:
       58
Query:
           FPKGSRCWVSGWGHTHPSHTYSSDMLQDTVVPLLSTQLCNSSCVYSGALTPRMLCAGYLD
```

```
Applicants:
           Guo
U.S.S.N.:
           09/981,151
                               | | | + | | + | + |
            1 |+ | |||| | |
                                                   |+| ||||| |+
           VPAGTTCTVSGWGRTSESSGSLPDTLQEVNVPIVSNATCRRAYSGGPAITDNMLCAGGLE 177
Sbjct: 118
          GRADACQGDSGGPLVCPDGDTWRLVGVVSWGR-GCAEPNHPGVYAKVAEFLDWI
Query:
             GGKDACQGDSGGPLVC-NDPRWVLVGIVSWGSYGCARPNKPGVYTRVSSYLDWI
      (of SEQ ID NO:27)
Query:
Sbjct: (SEQ ID NO:96)
```

At page 116, please replace Table 10I with the following:

Table 10I. Domain Analysis of NOV10

gnl|Pfam|pfam00089, trypsin, Trypsin. Proteins recognized include all proteins in families S1, S2A, S2B, S2C, and S5 in the classification of peptidases. Also included are proteins that are clearly members, but that lack peptidase activity, such as haptoglobin and protein Z (PRTZ*). (SEQ ID NO:97)

CD-Length = 217 residues, 100.0% aligned core = 211 bits (538), Expect = 6e-56

Query:	225	<pre>IVGGQSVAPGRWPWQASVALGFRHTCGGSVLAPRWVVTAAHCMHSFRLARLSSWRVHAGL +</pre>	284
Sbjct:	1	IVGGREAQAGSFPWQVSLQVSSGHFCGGSLISENWVLTAAHCVSGASSVRVVLGE	55
Query:	285	VSHSAVRPH-QGALVERIIPHPLYSAQNHDYDVALLRLQTALNFSDTVGAVCLPAKEQHF +	343
Sbjct:	56	HNLGTTEGTEQKFDVKKIIVHPNYNPDTNDIALLKLKSPVTLGDTVRPICLPSASSDL	113
Query:	344	PKGSRCWVSGWGHTHPSHTYSSDMLQDTVVPLLSTQLCNSSCVYSGALTPRMLCAGYLDG	403
Sbjct:	114	PVGTTCSVSGWGRTKNLGTSDTLQEVVVPIVSRETCRSAYGGTVTDTMICAGALGG	169
Query:	404	RADACQGDSGGPLVCPDGDTWRLVGVVSWGRGCAEPNHPGVYAKVAEFLDWI 455 +	
Sbjct:	170	K-DACQGDSGGPLVCSDGELVGIVSWGYGCAVGNYPGVYTRVSRYLDWI 217	
Query:	(of S	SEQ ID NO:27)	

Sbjct: (SEQ ID NO:97)